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110> University of California Carson, Dennis A. Corr, Maripat Rhee, Chae-Seo Lorenzo, Leoni M. Malini, Sen

<120> IMMUNOLOGIC COMPOSITIONS AND METHODS FOR STUDYING AND TREATING CANCERS EXPRESSING FRIZZLED ANTIGENS

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Val Ser Thr Phe Leu Ile Asp Met Glu Arg Phe Lys Tyr Pro Glu Arg 260 265 Pro Ile Ile Phe Leu Ser Ala Cys Tyr Leu Phe Val Ser Val Gly Tyr Leu Val Arq Leu Val Ala Gly His Glu Lys Val Ala Cys Ser Gly Gly 295 Ala Pro Gly Ala Gly Gly Arg Gly Gly Ala Gly Gly Ala Ala Ala Ala 310 315 Gly Ala Gly Ala Ala Gly Arg Gly Ala Ser Ser Pro Gly Ala Arg Gly 325 330 Glu Tyr Glu Glu Leu Gly Ala Val Glu Gln His Val Arg Tyr Glu Thr 345 Thr Gly Pro Ala Leu Cys Thr Val Val Phe Leu Leu Val Tyr Phe Phe 360 Gly Met Ala Ser Ser Ile Trp Trp Val Ile Leu Ser Leu Thr Trp Phe 375 Leu Ala Ala Gly Met Lys Trp Gly Asn Glu Ala Ile Ala Gly Tyr Ser 390 395 Gln Tyr Phe His Leu Ala Ala Trp Leu Val Pro Ser Val Lys Ser Ile 405 410 Ala Val Leu Ala Leu Ser Ser Val Asp Gly Asp Pro Val Ala Gly Ile 425 Cys Tyr Val Gly Asn Gln Ser Leu Asp Asn Leu Arg Gly Phe Val Leu 435 440 445 Ala Pro Leu Val Ile Tyr Leu Phe Ile Gly Thr Met Phe Leu Leu Ala 455 . . 460 Gly Phe Val Ser Leu Phe Arg Ile Arg Ser Val Ile Lys Gln Gly 470 475 Gly Pro Thr Lys Thr His Lys Leu Glu Lys Leu Met Ile Arg Leu Gly 490 Leu Phe Thr Val Leu Tyr Thr Val Pro Ala Ala Val Val Ala Cys 500 505 Leu Phe Tyr Glu Gln His Asn Arg Pro Arg Trp Glu Ala Thr His Asn 520 Cys Pro Cys Leu Arg Asp Leu Gln Pro Asp Gln Ala Arg Arg Pro Asp 535 540 Tyr Ala Val Phe Met Leu Lys Tyr Phe Met Cys Leu Val Val Gly Ile 550 555 Thr Ser Gly Val Trp Val Trp Ser Gly Lys Thr Leu Glu Ser Trp Arg 570 565 Ala Leu Cys Thr Arg Cys Cys Trp Ala Ser Lys Gly Ala Ala Val Gly 585 Ala Gly Ala Gly Gly Ser Gly 595

<210> 38

<211> 516

<212> PRT

<213> Homo sapiens

<400> 38

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 Cys Gln Glu Ile Thr Val Pro Met Cys Arg Gly Ile Gly Tyr Asn Leu 35
 40
 45

Thr His Met Pro Asn Gln Phe Asn His Asp Thr Gln Asp Glu Ala Gly Leu Glu Val His Gln Phe Trp Pro Leu Val Glu Ile Gln Cys Ser Pro Asp Leu Arg Phe Phe Leu Cys Thr Met Tyr Thr Pro Ile Cys Leu Pro Asp Tyr His Lys Pro Leu Pro Pro Cys Arg Ser Val Cys Glu Arg Ala Lys Ala Gly Cys Ser Pro Leu Met Arg Gln Tyr Gly Phe Ala Trp Pro Glu Arg Met Ser Cys Asp Arg Leu Pro Val Leu Gly Arg Asp Ala Glu Val Leu Cys Met Asp Tyr Asn Arg Ser Glu Ala Thr Thr Ala Pro Pro Arg Pro Phe Pro Ala Lys Pro Thr Leu Pro Gly Pro Pro Gly Ala Pro Ala Ser Gly Gly Arg Thr Gly Gln Val Pro Asn Cys Ala Val Pro Cys Tyr Gln Pro Ser Phe Ser Ala Asp Glu Arg Thr Phe Ala Thr Phe Trp Ile Gly Leu Trp Ser Val Leu Cys Phe Ile Ser Thr Ser Thr Thr Val Ala Thr Phe Leu Ile Asp Met Asp Thr Phe Arg Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Ala Cys Tyr Leu Cys Val Ser Leu Gly Phe Leu Val Arg Leu Val Val Gly His Ala Ser Val Ala Cys Ser Arg Glu His Asn His Ile His Tyr Glu Thr Thr Gly Pro Ala Leu Cys Thr Ile Val Phe Leu Leu Val Tyr Phe Phe Gly Met Ala Ser Ser Ile Trp Trp Val Ile Leu Ser Leu Thr Trp Phe Leu Ala Ala Met Lys Trp Gly Asn Glu Ala Ile Ala Gly Tyr Gly Gln Tyr Phe His Leu Ala Ala Trp Leu Ile Pro Ser Val Lys Ser Ile Thr Ala Leu Ala Leu Ser Ser Val Asp Gly Asp Pro Val Ala Gly Ile Cys Tyr Val Gly Asn Gln Asn Leu Asn Ser Leu Arg Arg Phe Val Leu Gly Pro Leu Val Leu Tyr Leu Leu Val Gly Thr Leu Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Ser Val Ile Lys Gln Gly Gly Thr Lys Thr Asp Lys Leu Glu Lys Leu Met Ile Arg Ile Gly Ile Phe Thr Leu Leu Tyr Thr Val Pro Ala Ser Ile Val Val Ala Cys Tyr Leu Tyr Glu Gln His Tyr Arg Glu Ser Trp Glu Ala Ala Leu Thr Cys Ala Cys Pro Gly His Asp Thr Gly Gln Pro Arg Ala Lys Pro Glu Tyr Trp Val Leu Met Leu Lys Tyr Phe Met Cys Leu Val Val Gly Ile Thr Ser Gly Val Trp Ile Trp Ser Gly Lys Thr Val Glu Ser Trp Arg Arg Phe Thr Ser Arg Cys Cys Cys Arg Pro Arg

500 505 510 Arg Gly His Lys 515 <210> 39 <211> 533 <212> PRT <213> Homo sapiens <400> 39 Met Ala Val Ala Pro Leu Arg Gly Ala Leu Leu Trp Gln Leu Leu 10 Ala Ala Gly Gly Ala Ala Leu Glu Ile Gly Arg Phe Asp Pro Glu Arg 25 Gly Arg Gly Ala Ala Pro Cys Gln Ala Val Glu Ile Pro Met Cys Arg Gly Ile Gly Tyr Asn Leu Thr Arg Met Pro Asn Leu Leu Gly His Thr 60 55 Ser Gln Gly Glu Ala Ala Ala Glu Leu Ala Glu Phe Ala Pro Leu Val 70 75 Gln Tyr Gly Cys His Ser His Leu Arg Phe Phe Leu Cys Ser Leu Tyr 85 90 Ala Pro Met Cys Thr Asp Gln Val Ser Thr Pro Ile Pro Ala Cys Arg 105 Pro Met Cys Glu Gln Ala Arg Leu Arg Cys Ala Pro Ile Met Glu Gln 120 125 Phe Asn Phe Gly Trp Pro Asp Ser Leu Asp Cys Ala Arg Leu Pro Thr 135 140 Arg Asn Asp Pro His Ala Leu Cys Met Glu Ala Pro Glu Asn Ala Thr 155 Ala Gly Pro Ala Glu Pro His Lys Gly Leu Gly Met Leu Pro Val Ala 165 170 Pro Arg Pro Ala Arg Pro Pro Gly Arg Ser Cys Ala Pro Arg Cys Gly 185 Pro Gly Val Glu Val Phe Trp Ser Arg Arg Asp Lys Asp Phe Ala Leu 195 200 205 Val Trp Met Ala Val Trp Ser Ala Leu Cys Phe Phe Ser Thr Ala Phe 215 220 Thr Val Leu Thr Phe Leu Leu Glu Pro His Arg Phe Gln Tyr Pro Glu 230 235 Arg Pro Ile Ile Phe Leu Ser Met Cys Tyr Asn Val Tyr Ser Leu Ala 250 245 Phe Leu Ile Arg Ala Val Ala Gly Ala Gln Ser Val Ala Cys Asp Gln 265 Glu Ala Gly Ala Leu Tyr Val Ile Gln Glu Gly Leu Glu Asn Thr Gly 275 280 Cys Thr Leu Val Phe Leu Leu Tyr Tyr Phe Gly Met Ala Ser Ser 295 Leu Trp Trp Val Val Leu Thr Leu Thr Trp Phe Leu Ala Ala Gly Lys 310 315 Lys Trp Gly His Glu Ala Ile Glu Ala His Gly Ser Tyr Phe His Met 325 330 Ala Ala Trp Gly Leu Pro Ala Leu Lys Thr Ile Val Ile Leu Thr Leu 345 Arg Lys Val Ala Gly Asp Glu Leu Thr Gly Leu Cys Tyr Val Ala Ser 360 365 Thr Asp Ala Ala Ala Leu Thr Gly Phe Val Leu Val Pro Leu Ser Gly

375 Tyr Leu Val Leu Gly Ser Ser Phe Leu Leu Thr Gly Phe Val Ala Leu 390 395 Phe His Ile Arg Lys Ile Met Lys Thr Gly Gly Thr Asn Thr Glu Lys 405 410 Leu Glu Lys Leu Met Val Lys Ile Gly Val Phe Ser Ile Leu Tyr Thr 425 Val Pro Ala Thr Cys Val Ile Val Cys Tyr Val Tyr Glu Arg Leu Asn 440 Met Asp Phe Trp Arg Leu Arg Ala Thr Glu Gln Pro Cys Ala Ala Ala 455 460 Ala Gly Pro Gly Gly Arg Arg Asp Cys Ser Leu Pro Gly Gly Ser Val 470 475 Pro Thr Val Ala Val Phe Met Leu Lys Ile Phe Met Ser Leu Val Val 485 490 Gly Ile Thr Ser Gly Val Trp Val Trp Ser Ser Lys Thr Phe Gln Thr 505 Trp Gln Ser Leu Cys Tyr Arg Lys Ile Ala Ala Gly Arg Ala Arg Ala 520 Lys Ala Cys Arg Ala 530 <210> 40 <211> 544 <212> PRT <213> Rat <400> 40 Leu Glu Ala Pro Leu Leu Gly Val Arg Ala Gln Pro Ala Gly Gln Val Ser Gly Pro Gly Gln Gln Arg Pro Pro Pro Gln Pro Gln Gln 25 Gly Gly Gln Gln Tyr Asn Gly Glu Arg Gly Ile Ser Ile Pro Asp His 40 Gly Tyr Cys Gln Pro Ile Ser Ile Pro Leu Cys Thr Asp Ile Ala Tyr 55 Asn Gln Thr Ile Met Pro Asn Leu Leu Gly His Thr Asn Gln Glu Asp 70 75 Ala Gly Leu Glu Val His Gln Phe Tyr Pro Leu Val Lys Val Gln Cys 85 90 Ser Ala Glu Leu Lys Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys 105 Thr Val Leu Glu Gln Ala Leu Pro Pro Cys Arg Ser Leu Cys Glu Arg 120 Ala Gln Gly Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln Trp Pro 135 140 Asp Thr Leu Lys Cys Glu Lys Phe Pro Val His Gly Ala Gly Glu Leu 150 155 Cys Val Gly Gln Asn Thr Ser Asp Lys Gly Thr Pro Thr Pro Ser Leu 165 170 Leu Pro Glu Phe Trp Thr Ser Asn Pro Gln His Gly Leu Gly Glu Lys 185 Asp Cys Gly Ala Pro Cys Glu Pro Thr Lys Val Tyr Gly Leu Met Tyr 200 Phe Gly Pro Glu Glu Leu Arg Phe Ser Arg Thr Trp Ile Gly Ile Trp 215 220 Ser Val Leu Cys Cys Ala Ser Thr Leu Phe Thr Val Leu Thr Tyr Leu

225 230 Val Asp Met Arg Arg Phe Ser Tyr Pro Glu Arg Pro Ile Ile Phe Leu 250 Ser Gly Cys Tyr Thr Ala Val Ala Val Ala Tyr Ile Ala Gly Phe Leu 265 Leu Glu Asp Arg Val Val Cys Asn Asp Lys Phe Ala Glu Asp Gly Ala 280 Arg Thr Val Ala Gln Gly Thr Lys Lys Glu Gly Cys Thr Ile Leu Phe 295 300 Met Met Leu Tyr Phe Phe Ser Met Ala Ser Ser Ile Trp Trp Val Ile 310 315 Leu Ser Leu Thr Trp Phe Leu Ala Ala Gly Met Lys Trp Gly His Glu 330 Ala Ile Glu Ala Asn Ser Gln Tyr Phe His Leu Ala Ala Trp Ala Val 345 Pro Ala Ile Lys Thr Ile Thr Ile Leu Ala Leu Gly Gln Val Asp Gly 360 Asp Val Leu Ser Gly Val Cys Phe Val Gly Leu Asn Asn Val Asp Ala 375 380 Leu Arg Gly Phe Val Leu Ala Pro Leu Phe Val Tyr Leu Phe Ile Gly 390 395 Thr Ser Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Thr 405 410 Ile Met Lys His Asp Gly Thr Lys Thr Glu Lys Leu Glu Lys Leu Met 425 420 430 Val Arg Ile Gly Val Phe Ser Val Leu Tyr Thr Val Pro Ala Thr Ile 440 Val Ile Ala Cys Tyr Phe Tyr Glu Gln Ala Phe Arg Asp Gln Trp Glu 455 460 Arg Ser Trp Val Ala Gln Ser Cys Lys Ser Tyr Ala Ile Pro Cys Pro 475 470 His Leu Gln Gly Gly Gly Val Pro Pro His Pro Pro Met Ser Pro 485 490 Asp Phe Thr Val Phe Met Ile Lys Tyr Leu Met Thr Leu Ile Val Gly 505 Ile Thr Ser Gly Phe Trp Ile Trp Ser Gly Lys Thr Leu Asn Ser Trp 520 Arg Lys Phe Tyr Thr Arg Leu Thr Asn Ser Lys Gln Gly Glu Thr Thr 530 535

<210> 41

<211> 529

<212> PRT

<213> Rat

<400> 41

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Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val Leu Glu Gln Ala Ile Pro Pro Cys Arg Ser Ile Cys Glu Arg Ala Arg Gln Gly Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln Trp Pro Glu Arg Leu Arg Cys Glu His Phe Pro Arg His Gly Ala Glu Gln Ile Cys Val Gly Gln Asn His Ser Glu Asp Gly Thr Pro Ala Leu Leu Thr Thr Ala Pro Pro Ser Gly Leu Gln Pro Gly Leu Gly Glu Arg Asp Cys Ala Ala Pro Cys Glu Pro Ala Arg Pro Asp Gly Ser Met Phe Phe Ser His His His Thr Arg Phe Ala Arg Leu Trp Ile Leu Thr Trp Ser Val Leu Cys Cys Ala Ser Thr Phe Phe Thr Val Thr Thr Ser Leu Val Ala Met Gln Arg Phe Arg Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Gly Cys Tyr Thr Met Val Ser Val Ala Tyr Ile Ala Gly Phe Val Leu Gln Glu Arg Val Val Cys Asn Glu Arg Phe Ser Glu Asp Gly Tyr Arg Thr Val Gly Gln Gly Thr Lys Lys Glu Gly Cys Thr Ile Leu Phe Met Met Leu Tyr Phe Phe Ser Met Ala Ser Ser Ile Trp Trp Val Ile Leu Ser Leu Thr Trp Phe Leu Ala Ala Gly Met Lys Trp Gly His Ala Ala Ile Glu Ala Asn Ser Gln Tyr Phe His Leu Ala Ala Trp Ala Val Pro Ala Val Lys Thr Ile Thr Ile Leu Ala Met Gly Gln Ile Asp Gly Asp Leu Leu Ser Gly Val Cys Phe Val Gly Leu Asn Arg Leu Asp Pro Leu Arg Gly Phe Val Leu Ala Pro Leu Phe Val Tyr Leu Phe Ile Gly Thr Ser Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Thr Ile Met Lys His Asp Gly Thr Lys Thr Glu Pro Leu Glu Arg Leu Met Val Arg Ile Gly Val Phe Ser Val Leu Tyr Thr Val Pro Ala Thr Ile Val Ile Ala Cys Tyr Phe Tyr Glu Gln Ala Phe Arg Glu His Trp Glu Arg Ser Trp Val Ser Gln His Cys Lys Ser Leu Ala Ile Pro Cys Pro Ala His Tyr Thr Pro Arg Thr Ser Pro Asp Phe Thr Val Tyr Met Ile Lys Tyr Leu Met Thr Leu Ile Val Gly Ile Thr Ser Gly Phe Trp Ile Trp Ser Gly Lys Thr Leu His Ser Trp Arg Lys Phe Tyr Thr Arg Leu Thr Asn Ser Arg His Gly Glu Thr Thr

<210> 42 <211> 536 <212> PRT <213> Drosophila <400> 42 Ile Leu Pro Thr

Ile Leu Pro Thr Leu Ile Gln Gly Val Gln Arg Tyr Asp Gln Ser Pro Leu Asp Ala Ser Pro Tyr Tyr Arg Ser Gly Gly Leu Met Ala Ser 20 25 Ser Gly Thr Glu Leu Asp Gly Leu Pro His His Asn Arg Cys Glu Pro Ile Thr Ile Ser Ile Cys Lys Asn Ile Pro Tyr Asn Met Thr Ile Met Pro Asn Leu Ile Gly His Thr Lys Gln Glu Glu Ala Gly Leu Glu Val His Gln Phe Ala Pro Leu Val Lys Ile Gly Cys Ser Asp Asp Leu Gln 85 Leu Phe Leu Cys Ser Leu Tyr Val Pro Val Cys Thr Ile Leu Glu Arg 105 Pro Ile Pro Pro Cys Arg Ser Leu Cys Glu Ser Ala Arg Val Cys Glu 120 Lys Leu Met Lys Thr Tyr Asn Phe Asn Trp Pro Glu Asn Leu Glu Cys 135 140 Ser Lys Phe Pro Val His Gly Glu Asp Leu Cys Val Ala Glu Asn 155 Thr Thr Ser Ser Ala Ser Thr Ala Ala Thr Pro Thr Arg Ser Val Ala 165 170 Val Gly Gly Lys Asp Leu His Asp Cys Gly Ala Pro Cys His Ala Met 185 190 Phe Phe Pro Glu Arg Glu Arg Thr Val Leu Arg Tyr Trp Val Gly Ser 200 205 Trp Ala Ala Val Cys Val Ala Ser Cys Leu Phe Thr Val Leu Thr Phe 215 220 Leu Ile Asp Ser Ser Arg Phe Arg Tyr Pro Glu Arg Ala Ile Val Phe 235 Leu Ala Val Cys Tyr Leu Val Val Gly Cys Ala Tyr Val Ala Gly Leu 250 245 Gly Ala Gly Asp Ser Val Ser Cys Arg Glu Pro Phe Pro Pro Pro Val 265 Lys Leu Gly Arg Leu Gln Met Met Ser Thr Ile Thr Gln Gly His Arg 280 Gln Thr Thr Ser Cys Thr Val Leu Phe Met Ala Leu Tyr Phe Cys Cys 295 300 Met Ala Ala Phe Ala Trp Trp Ser Cys Leu Ala Phe Ala Trp Phe Leu 310 315 Ala Ala Gly Leu Lys Trp Gly His Glu Ala Ile Glu Asn Lys Ser His 325 330 Leu Phe His Leu Val Ala Trp Ala Val Pro Ala Leu Gln Thr Ile Ser 345 Val Leu Ala Leu Ala Lys Val Glu Gly Asp Ile Leu Ser Gly Val Cys 360 365 Phe Val Gly Gln Leu Asp Thr His Ser Leu Gly Ala Phe Leu Ile Leu 375 380 Pro Leu Cys Ile Tyr Leu Ser Ile Gly Ala Leu Phe Leu Leu Ala Gly 395 390 400

Phe Ile Ser Leu Phe Arg Ile Arg Thr Val Met Lys Thr Asp Gly Lys 410 415 Arg Thr Asp Lys Leu Glu Arg Leu Met Leu Arg Ile Gly Phe Phe Ser 420 425 Gly Leu Phe Ile Leu Pro Ala Val Gly Leu Leu Gly Cys Leu Phe Tyr 440 Glu Tyr Tyr Asn Phe Asp Glu Trp Met Ile Gln Trp His Arg Asp Ile 455 Cys Lys Pro Phe Ser Ile Pro Cys Pro Ala Ala Arg Ala Pro Gly Ser 470 475 Pro Glu Ala Arg Pro Ile Phe Gln Ile Phe Met Val Lys Tyr Leu Cys 485 490 Ser Met Leu Val Gly Val Thr Ser Ser Val Trp Leu Tyr Ser Ser Lys 505 Thr Met Val Ser Trp Arg Asn Phe Val Glu Arg Leu Gln Gly Lys Glu 520 Pro Arg Thr Arg Ala Gln Ala Tyr 530

<210> 43 <211> 570 <212> PRT

<213> Drosophila

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165

Ser Gly Gly Lys Arg Lys Gln Gly Gly Ser Gly Ser Gly Gly Ser Gly 195

Ala Gly Gly Ser Ser Gly Ser Thr Ser Thr Lys Pro Cys Arg Gly Arg 210

Gln Arg Ile Ala Gly Val Pro Asn Cys Gly Ile Pro Cys Lys Gly Pro 225

Phe Phe Ser Asn Asp Glu Lys Asp Phe Ala Gly Leu Trp Ile Ala Leu 245

Asp Pro Asp Asn Leu Cys Met Glu Gln Pro Ser Tyr Thr Glu Ala Gly

Ser Gly Gly Ser Ser Gly Gly Ser Gly Ser Gly Ser Gly 180 185 190

170

Trp Ser Gly Leu Cys Phe Cys Ser Thr Leu Met Thr Leu Thr Thr Phe 265 Ile Ile Asp Thr Glu Arg Phe Lys Tyr Pro Glu Arg Pro Ile Val Phe 280 285 275 Leu Ser Ala Cys Tyr Phe Met Val Ala Val Gly Tyr Leu Ser Arg Asn 295 Phe Leu Gln Asn Glu Glu Ile Ala Cys Asp Gly Leu Leu Leu Arg Glu 310 315 Ser Ser Thr Gly Pro His Ser Cys Thr Leu Val Phe Leu Leu Thr Tyr 330 325 Phe Phe Gly Met Ala Ser Ser Ile Trp Trp Val Ile Leu Thr Phe Thr 345 Trp Phe Leu Ala Ala Gly Leu Lys Trp Gly Asn Glu Ala Ile Thr Lys 360 His Ser Gln Tyr Phe His Leu Ala Ala Trp Leu Ile Pro Thr Val Gln 375 Ser Val Ala Val Leu Leu Ser Ala Val Asp Gly Asp Pro Ile Leu 390 395 Gly Ile Cys Tyr Val Gly Asn Leu Asn Pro Asp His Leu Lys Thr Phe 410 405 Val Leu Ala Pro Leu Phe Val Tyr Leu Val Ile Gly Thr Thr Phe Leu 425 Met Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Ser Val Ile Lys Gln 440 445 435 Gln Gly Gly Val Gly Ala Gly Val Lys Ala Asp Lys Leu Glu Lys Leu 455 Met Ile Arg Ile Gly Ile Phe Ser Val Leu Tyr Thr Val Pro Ala Thr 475 470 Ile Val Ile Gly Cys Tyr Leu Tyr Glu Ala Ala Tyr Phe Glu Asp Trp 490 485 Ile Lys Ala Leu Ala Cys Pro Cys Ala Gln Val Lys Gly Pro Gly Lys 505 Lys Pro Leu Tyr Ser Val Leu Met Leu Lys Tyr Phe Met Ala Leu Ala 520 Val Gly Ile Thr Ser Gly Val Trp Ile Trp Ser Gly Lys Thr Leu Glu 535 Ser Trp Arg Arg Phe Trp Arg Leu Leu Gly Ala Pro Asp Arg Thr 550 555 Gly Ala Asn Gln Ala Leu Ile Lys Gln Arg 565

<210> 44

<211> 647

<212> PRT

<213> Homo sapiens

<400> 44

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Gln Gly Pro Gly Pro Gly Gln Gln Pro Pro Pro Pro Gln Gln Gln Gln Ser Gly Gln Gln Tyr Asn Gly Glu Arg Gly Ile Ser Val Pro Asp 105 His Gly Tyr Cys Gln Pro Ile Ser Ile Pro Leu Cys Thr Asp Ile Ala 120 125 Tyr Asn Gln Thr Ile Met Pro Asn Leu Leu Gly His Thr Asn Gln Glu 135 Asp Ala Gly Leu Glu Val His Gln Phe Tyr Pro Leu Val Lys Val Gln 150 155 Cys Ser Ala Glu Leu Lys Phe Phe Leu Cys Ser Met Tyr Ala Pro Val 165 170 Cys Thr Val Leu Glu Gln Ala Leu Pro Pro Cys Arg Ser Leu Cys Glu 185 Arg Ala Arg Gln Gly Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln 205 200 Trp Pro Asp Thr Leu Lys Cys Glu Lys Phe Pro Val His Gly Ala Gly 215 220 Glu Leu Cys Val Gly Gln Asn Thr Ser Asp Lys Gly Thr Pro Thr Pro 230 235 Ser Leu Leu Pro Glu Phe Trp Thr Ser Asn Pro Gln His Gly Gly Gly 245 250 Gly His Arg Gly Gly Phe Pro Gly Gly Ala Gly Ala Ser Glu Arg Gly 265 Lys Phe Ser Cys Pro Arg Ala Leu Lys Val Pro Ser Tyr Leu Asn Tyr 285 275 280 His Phe Leu Gly Glu Lys Asp Cys Gly Ala Pro Cys Glu Pro Thr Lys 295 Val Tyr Gly Leu Met Tyr Phe Gly Pro Glu Glu Leu Arg Phe Ser Arg Thr Trp Ile Gly Ile Trp Ser Val Leu Cys Cys Ala Ser Thr Leu Phe 325 330 Thr Val Leu Thr Tyr Leu Val Asp Met Arg Arg Phe Ser Tyr Pro Glu 345 Arg Pro Ile Ile Phe Leu Ser Gly Cys Tyr Thr Ala Val Ala Val Ala 360 Tyr Ile Ala Gly Phe Leu Leu Glu Asp Arg Val Val Cys Asn Asp Lys 375 Phe Ala Glu Asp Gly Ala Arg Thr Val Ala Gln Gly Thr Lys Lys Glu 390 395 Gly Cys Thr Ile Leu Phe Met Met Leu Tyr Phe Phe Ser Met Ala Ser 405 410 Ser Ile Trp Trp Val Ile Leu Ser Leu Thr Trp Phe Leu Ala Ala Gly 425 Met Lys Trp Gly His Glu Ala Ile Glu Ala Asn Ser Gln Tyr Phe His 440 Leu Ala Ala Trp Ala Val Pro Ala Ile Lys Thr Ile Thr Ile Leu Ala 455 Leu Gly Gln Val Asp Gly Asp Val Leu Ser Gly Val Cys Phe Val Gly 470 . 475 Leu Asn Asn Val Asp Ala Leu Arg Gly Phe Val Leu Ala Pro Leu Phe 485 490 Val Tyr Leu Phe Ile Gly Thr Ser Phe Leu Leu Ala Gly Phe Val Ser 505 Leu Phe Arg Ile Arg Thr Ile Met Lys His Asp Gly Thr Lys Thr Glu 520 Lys Leu Glu Lys Leu Met Val Arg Ile Gly Val Phe Ser Val Leu Tyr

535 540 Thr Val Pro Ala Thr Ile Val Ile Ala Cys Tyr Phe Tyr Glu Gln Ala 550 555 Phe Arg Asp Gln Trp Glu Arg Ser Trp Val Ala Gln Ser Cys Lys Ser 565 570 Tyr Ala Ile Pro Cys Pro His Leu Gln Ala Gly Gly Gly Ala Pro Pro 585 His Pro Pro Met Ser Pro Asp Phe Thr Val Phe Met Ile Lys Tyr Leu 600 Met Thr Leu Ile Val Gly Ile Thr Ser Gly Phe Trp Ile Trp Ser Gly 615 620 Lys Thr Leu Asn Ser Trp Arg Lys Phe Tyr Thr Arg Leu Thr Asn Ser 630 635 Lys Gln Gly Glu Thr Thr Val 645

<210> 45 <211> 626 <212> PRT <213> Mouse

<400> 45

Met Ala Glu Glu Ala Ala Pro Ser Glu Ser Arg Ala Ala Gly Arg Leu Ser Leu Glu Leu Cys Ala Glu Ala Leu Pro Gly Arg Arg Glu Glu Val 20 25 Gly His Glu Asp Thr Ala Ser His Arg Arg Pro Arg Ala Asp Pro Arg Arg Trp Ala Ser Gly Leu Leu Leu Leu Trp Leu Leu Glu Ala Pro Leu Leu Gly Val Arg Ala Gln Ala Gly Gln Val Ser Gly Pro 70 75 Gly Gln Gln Ala Pro Pro Pro Gln Pro Gln Gln Ser Gly Gln Gln Tyr Asn Gly Glu Arg Gly Ile Ser Ile Pro Asp His Gly Tyr Cys Gln 105 Pro Ile Ser Ile Pro Leu Cys Thr Asp Met Ala Tyr Asn Gln Thr Ile 120 Met Pro Asn Leu Leu Gly His Thr Asn Gln Glu Asp Ala Gly Leu Glu 135 140 Val His Gln Phe Tyr Pro Leu Val Lys Val Gln Cys Ser Ala Glu Leu 150 155 Lys Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val Leu Glu 170 Gln Ala Leu Pro Pro Cys Arg Ser Leu Cys Glu Arg Ala Arg Gln Gly 180 185 190 Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln Trp Pro Asp Thr Leu 200 Lys Cys Glu Lys Phe Pro Val His Gly Ala Gly Glu Leu Cys Val Gly 215 220 Gln Asn Thr Ser Asp Lys Gly Thr Pro Thr Pro Ser Leu Leu Pro Glu 230 235 Phe Trp Thr Ser Asn Gly Gln His Gly Gly Gly Tyr Arg Gly Gly 245 250 Tyr Pro Gly Gly Ala Gly Thr Val Glu Arg Gly Lys Phe Ser Cys Pro 265 Arg Ala Leu Arg Val Pro Ser Tyr Leu Asn Tyr His Phe Leu Gly Glu

275 280 285 Lys Asp Cys Gly Ala Pro Cys Glu Pro Thr Lys Val Tyr Gly Leu Met 295 Tyr Phe Gly Pro Glu Glu Leu Arg Phe Ser Arg Thr Trp Ile Gly Ile 315 310 Trp Ser Val Leu Cys Cys Ala Ser Thr Leu Phe Thr Val Leu Thr Tyr 325 330 Leu Val Asp Met Pro Arg Phe Ser Tyr Pro Glu Arg Pro Ile Ile Ser 345 Leu Ser Gly Cys Tyr Thr Ala Val Ala Val Ala Tyr Ile Ala Gly Phe 365 360 Leu Leu Glu Asp Arg Val Val Cys Asn Asp Lys Phe Ala Glu Asp Gly 375 Ala Arg Thr Val Ala Gln Gly Thr Asn Lys Glu Gly Cys Thr Ile Leu 390 395 Phe Met Met Leu Tyr Phe Phe Ser Met Ala Ser Ser Ile Trp Trp Val 405 410 Ile Leu Ser Leu Thr Trp Phe Leu Ala Ala Gly Met Lys Trp Gly His 425 Glu Ala Ile Glu Ala Asn Ser Gln Tyr Phe His Leu Ala Ala Trp Ala 440 Val Pro Ala Ile Lys Thr Ile Thr Ile Leu Ala Leu Gly Gln Val Asp 455 Gly Asp Val Leu Ser Gly Val Cys Phe Leu Gly Leu Asn Asn Val Asp 470 475 Ala Leu Arg Gly Phe Val Leu Ala Pro Leu Phe Val Tyr Leu Phe Ile 490 Gly Thr Ser Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile Arg 505 Thr Ile Met Lys His Asp Gly Thr Lys Thr Glu Lys Leu Glu Lys Leu 525 520 Met Val Arg Ile Gly Val Phe Ser Val Leu Tyr Thr Val Pro Ala Thr 535 540 Ile Val Ile Ala Cys Tyr Phe Tyr Glu Gln Ala Phe Arg Asp Gln Trp 550 555 Glu Arg Ser Trp Val Ala Gln Ser Cys Lys Ser Tyr Ala Ile Pro Cys 570 Pro His Leu Gln Gly Gly Gly Val Pro Pro His Pro Pro Met Ser 585 Pro Asp Phe Thr Val Phe Met Ile Lys Tyr Leu Met Thr Leu Asn Ser 600 Trp Arg Lys Phe Tyr Thr Arg Leu Thr Asn Ser Lys Gln Gly Glu Thr 610 615 Thr Val 625 <210> 46 <211> 565 <212> PRT <213> Homo sapiens <400> 46 Met Arg Pro Arg Ser Ala Leu Pro Arg Leu Leu Pro Leu Leu Leu Leu Pro Ala Ala Gly Pro Ala Gln Phe His Gly Glu Lys Gly Ile Ser

Ile Pro Asp His Gly Phe Cys Gln Pro Ile Ser Ile Pro Leu Cys Thr

40 Asp Ile Ala Tyr Asn Gln Thr Ile Met Pro Asn Leu Leu Gly His Thr Asn Gln Glu Asp Ala Gly Leu Glu Val His Gln Phe Tyr Pro Leu Val 70 Lys Val Gln Cys Ser Pro Glu Leu Arg Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val Leu Glu Gln Ala Ile Pro Pro Cys Arg Ser 105 Ile Cys Glu Arg Ala Arg Gln Gly Cys Glu Ala Leu Met Asn Lys Phe 120 125 Gly Phe Gln Trp Pro Glu Arg Leu Arg Cys Glu His Phe Pro Arg His 135 Gly Ala Glu Gln Ile Cys Val Gly Gln Asn His Ser Glu Asp Gly Ala 155 Pro Ala Leu Leu Thr Thr Ala Pro Pro Pro Gly Leu Gln Pro Gly Ala 165 Gly Gly Thr Pro Gly Gly Pro Gly Gly Gly Gly Ala Pro Pro Arg Tyr 185 Ala Thr Leu Glu His Pro Phe His Cys Pro Arg Val Leu Lys Val Pro 200 Ser Tyr Leu Ser Tyr Lys Phe Leu Gly Glu Arg Asp Cys Ala Ala Pro 215 Cys Glu Pro Ala Arg Pro Asp Gly Ser Met Phe Phe Ser Gln Glu Glu 230 235 Thr Arg Phe Ala Arg Leu Trp Ile Leu Thr Trp Ser Val Leu Cys Cys 245 250 Ala Ser Thr Phe Phe Thr Val Thr Tyr Leu Val Asp Met Gln Arg 265 Phe Arg Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Gly Cys Tyr Thr 280 Met Val Ser Val Ala Tyr Ile Ala Gly Phe Val Leu Gln Glu Arg Val 295 300 Val Cys Asn Glu Arg Phe Ser Glu Asp Gly Tyr Arg Thr Val Val Gln 310 315 Gly Thr Lys Lys Glu Gly Cys Thr Ile Leu Phe Met Met Leu Tyr Phe 325 330 Phe Ser Met Ala Ser Ser Ile Trp Trp Val Ile Leu Ser Leu Thr Trp 345 Phe Leu Ala Ala Gly Met Lys Trp Gly His Glu Ala Ile Glu Ala Asn 360 Ser Gln Tyr Phe His Leu Ala Ala Trp Ala Val Pro Ala Val Lys Thr 375 Ile Thr Ile Leu Ala Met Gly Gln Ile Asp Gly Asp Leu Leu Ser Gly 390 395 Val Cys Phe Val Gly Leu Asn Ser Leu Asp Pro Leu Arg Gly Phe Val 405 410 Leu Ala Pro Leu Phe Val Tyr Leu Phe Ile Gly Thr Ser Phe Leu Leu 425 Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Thr Ile Met Lys His Asp 440 Gly Thr Lys Thr Glu Lys Leu Glu Arg Leu Met Val Arg Ile Gly Val 455 460 Phe Ser Val Leu Tyr Thr Val Pro Ala Thr Ile Val Ile Ala Cys Tyr 470 475 Phe Tyr Glu Gln Ala Phe Arg Glu His Trp Glu Arg Ser Trp Val Ser 490 485

<210> 47 <211> 666 <212> PRT <213> Homo sapiens

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Ser Glu Ala Ile Glu Lys Lys Ala Leu Leu Phe His Ala Ser Ala Trp 330 Gly Ile Pro Gly Thr Leu Thr Ile Ile Leu Leu Ala Met Asn Lys Ile 345 Glu Gly Asp Asn Ile Ser Gly Val Cys Phe Val Gly Leu Tyr Asp Val 360 Asp Ala Leu Arg Tyr Phe Val Leu Ala Pro Leu Cys Leu Tyr Val Val 375 Val Gly Val Ser Leu Leu Leu Ala Gly Ile Ile Ser Leu Asn Arg Val 390. 395 Arg Ile Glu Ile Pro Leu Glu Lys Glu Asn Gln Asp Lys Leu Val Lys 405 410 Phe Met Ile Arg Ile Gly Val Phe Ser Ile Leu Tyr Leu Val Pro Leu 425 Leu Val Val Ile Gly Cys Tyr Phe Tyr Glu Gln Ala Tyr Arg Gly Ile 440 Trp Glu Thr Trp Ile Gln Glu Arg Cys Arg Glu Tyr His Ile Pro 455 460 Cys Pro Tyr Gln Val Thr Gln Met Ser Arg Pro Asp Leu Ile Leu Phe 470 475 Leu Met Lys Tyr Leu Met Ala Leu Ile Val Gly Ile Pro Ser Val Phe 485 490 Trp Val Gly Ser Lys Lys Thr Cys Phe Glu Trp Ala Ser Phe Phe His 505 Gly Arg Arg Lys Lys Glu Ile Val Asn Glu Ser Arg Gln Val Leu Gln 520 525 Glu Pro Asp Phe Ala Gln Ser Leu Leu Arg Asp Pro Asn Thr Pro Ile 535 540 Ile Arg Lys Ser Arg Gly Thr Ser Thr Gln Gly Thr Ser Thr His Ala 550 555 Ser Ser Thr Gln Leu Ala Met Val Asp Asp Gln Arg Ser Lys Ala Gly 565 570 Ser Ile His Ser Lys Val Ser Ser Tyr His Gly Ser Leu His Arg Ser 585 Arg Asp Gly Arg Tyr Thr Pro Cys Ser Tyr Arg Gly Met Glu Glu Arg 600 Leu Pro His Gly Ser Met Ser Arg Leu Thr Asp His Ser Arg His Ser 615 620 Ser Ser His Arg Leu Asn Glu Gln Ser Arg His Ser Ser Ile Arg Asp 630 635 Leu Ser Asn Asn Pro Met Thr His Ile Thr His Gly Thr Ser Met Asn 645 650 Arg Val Ile Glu Glu Asp Gly Thr Ser Ala 660

<210> 48

<211> 666

<212> PRT

<213> Mouse

<400> 48

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 Asp Leu Trp Leu Leu Thr Val Phe

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 20
 25
 30

 Leu Arg Met Cys Gln Asp Leu Pro Tyr Asn Thr Thr Phe Met Pro Asn
 35
 40
 45

Leu Leu Asn His Tyr Asp Gln Gln Thr Ala Ala Leu Ala Met Glu Pro Phe His Pro Met Val Asn Leu Asp Cys Ser Arg Asp Phe Arg Pro Phe Leu Cys Ala Leu Tyr Ala Pro Ile Cys Met Glu Tyr Gly Arg Val Thr Leu Pro Cys Arg Arg Leu Cys Gln Arg Ala Tyr Ser Glu Cys Ser Lys Leu Met Glu Met Phe Gly Val Pro Trp Pro Glu Asp Met Glu Cys Ser Arg Phe Pro Asp Cys Asp Glu Pro Tyr Pro Arg Leu Val Asp Leu Asn Leu Val Gly Asp Pro Thr Glu Gly Ala Pro Val Ala Val Gln Arg Asp Tyr Gly Phe Trp Cys Pro Arg Glu Leu Lys Ile Asp Pro Asp Leu Gly Tyr Ser Phe Leu His Val Arg Asp Cys Ser Pro Pro Cys Pro Asn Met Tyr Phe Arg Arg Glu Glu Leu Ser Phe Ala Arg Tyr Phe Ile Gly Leu Ile Ser Ile Ile Cys Leu Ser Ala Thr Leu Phe Thr Phe Leu Thr Phe Leu Ile Asp Val Thr Arg Phe Arg Tyr Pro Glu Arg Pro Ile Ile Phe Tyr Ala Val Cys Tyr Met Met Val Ser Leu Ile Phe Phe Ile Gly Phe Leu Leu Glu Asp Arg Val Ala Cys Asn Ala Ser Ser Pro Ala Gln Tyr Lys Ala Ser Thr Val Thr Gln Gly Ser His Asn Lys Ala Cys Thr Met Leu Phe Met Val Leu Tyr Phe Phe Thr Met Ala Gly Ser Val Trp Trp Val Ile Leu Thr Ile Thr Trp Phe Leu Ala Ala Val Pro Lys Trp Gly Ser Glu Ala Ile Glu Lys Lys Ala Leu Leu Phe His Ala Ser Ala Trp Gly Ile Pro Gly Thr Leu Thr Ile Ile Leu Leu Ala Met Asn Lys Ile Glu Gly Asp Asn Ile Ser Gly Val Cys Phe Val Gly Leu Tyr Asp Val Asp Ala Leu Arg Tyr Phe Val Leu Ala Pro Leu Cys Leu Tyr Val Val Val Gly Val Ser Leu Leu Leu Ala Gly Ile Ile Ser Leu Asn Arg Val Arg Ile Glu Ile Pro Leu Glu Lys Glu Asn Gln Asp Lys Leu Val Lys Phe Met Ile Arg Ile Gly Val Phe Ser Ile Leu Tyr Leu Val Pro Leu Leu Val Val Ile Gly Cys Tyr Phe Tyr Glu Gln Ala Tyr Arg Gly Ile Trp Glu Thr Trp Ile Gln Glu Arg Cys Arg Glu Tyr His Ile Pro Cys Pro Tyr Gln Val Thr Gln Met Ser Arg Pro Asp Leu Ile Leu Phe Leu Met Lys Tyr Leu Met Ala Leu Ile Val Gly Ile Pro Ser Ile Phe Trp Val Gly Ser Lys Lys Thr Cys Phe Glu Trp Ala Ser Phe Phe His

500 505 Gly Arg Arg Lys Lys Glu Ile Val Asn Glu Ser Arg Gln Val Leu Gln 520 Glu Pro Asp Phe Ala Gln Ser Leu Leu Arg Asp Pro Asn Thr Pro Ile 535 540 Ile Arg Lys Ser Arg Gly Thr Ser Thr Gln Gly Thr Ser Thr His Ala 550 555 Ser Ser Thr Gln Leu Ala Met Val Asp Asp Gln Arg Ser Lys Ala Gly 565 570 Ser Val His Ser Lys Val Ser Ser Tyr His Gly Ser Leu His Arg Ser 585 590 Arg Asp Gly Arg Tyr Thr Pro Cys Ser Tyr Arg Gly Met Glu Glu Arg 600 Leu Pro His Gly Ser Met Ser Arg Leu Thr Asp His Ser Arg His Ser 615 620 Ser Ser His Arg Leu Asn Glu Gln Ser Arg His Ser Ser Ile Arg Asp 630 635 Leu Ser Asn Asn Pro Met Thr His Ile Thr His Gly Thr Ser Met Asn 645 650 Arg Val Ile Glu Glu Asp Gly Thr Ser Ala

<210> 49

<211> 537

<212> PRT

<213> Homo sapiens

<400> 49

Met Ala Trp Arg Gly Ala Gly Pro Ser Val Pro Gly Ala Pro Gly Gly Val Gly Leu Ser Leu Gly Leu Leu Leu Gln Leu Leu Leu Leu Gly 25 Pro Ala Arg Gly Phe Gly Asp Glu Glu Glu Arg Arg Cys Asp Pro Ile 40 Arg Ile Ser Met Cys Gln Asn Leu Gly Tyr Asn Val Thr Lys Met Pro 55 Asn Leu Val Gly His Glu Leu Gln Thr Asp Ala Glu Leu Gln Leu Thr 70 75 Thr Phe Thr Pro Leu Ile Gln Tyr Gly Cys Ser Ser Gln Leu Gln Phe 85 90 Phe Leu Cys Ser Val Tyr Val Pro Met Cys Thr Glu Lys Ile Asn Ile 105 Pro Ile Gly Pro Cys Gly Gly Met Cys Leu Ser Val Lys Arg Arg Cys Glu Pro Val Leu Lys Glu Phe Gly Phe Ala Trp Pro Glu Ser Leu Asn 135 140 Cys Ser Lys Phe Pro Pro Gln Asn Asp His Asn His Met Cys Met Glu 150 155 Gly Pro Gly Asp Glu Glu Val Pro Leu Pro His Lys Thr Pro Ile Gln 165 170 Pro Gly Glu Glu Cys His Ser Val Gly Thr Asn Ser Asp Gln Tyr Ile 185 . Trp Val Lys Arg Ser Leu Asn Cys Val Leu Lys Cys Gly Tyr Asp Ala 200 Gly Leu Tyr Ser Arg Ser Ala Lys Glu Phe Thr Asp Ile Trp Met Ala 215 Val Trp Ala Ser Leu Cys Phe Ile Ser Thr Ala Phe Thr Val Leu Thr

235 230 Phe Leu Ile Asp Ser Ser Arg Phe Ser Tyr Pro Glu Arg Pro Ile Ile 250 Phe Leu Ser Met Cys Tyr Asn Ile Tyr Ser Ile Ala Tyr Ile Val Arg 265 Leu Thr Val Gly Arg Glu Arg Ile Ser Cys Asp Phe Glu Glu Ala Ala 280 Glu Pro Val Leu Ile Gln Glu Gly Leu Lys Asn Thr Gly Cys Ala Ile 295 300 Ile Phe Leu Leu Met Tyr Phe Phe Gly Met Ala Ser Ser Ile Trp Trp 310 315 Val Ile Leu Thr Leu Thr Trp Phe Leu Ala Ala Gly Leu Lys Trp Gly 330 His Glu Ala Ile Glu Met His Ser Ser Tyr Phe His Ile Ala Ala Trp 345 Ala Ile Pro Ala Val Lys Thr Ile Val Ile Leu Ile Met Arg Leu Val 360 Asp Ala Asp Glu Leu Thr Gly Leu Cys Tyr Val Gly Asn Gln Asn Leu 375 380 Asp Ala Leu Thr Gly Phe Val Val Ala Pro Leu Phe Thr Tyr Leu Val 390 395 Ile Gly Thr Leu Phe Ile Ala Ala Gly Leu Val Ala Leu Phe Lys Ile 405 410 Arg Ser Asn Leu Gln Lys Asp Gly Thr Lys Thr Asp Lys Leu Glu Arg 430 425 420 Leu Met Val Lys Ile Gly Val Phe Ser Val Leu Tyr Thr Val Pro Ala Thr Cys Val Ile Ala Cys Tyr Phe Tyr Glu Ile Ser Asn Trp Ala Leu 455 460 Phe Arg Tyr Ser Ala Asp Asp Ser Asn Met Ala Val Glu Met Leu Lys 470 475 Ile Phe Met Ser Leu Leu Val Gly Ile Thr Ser Gly Met Trp Ile Trp 485 490 Ser Ala Lys Thr Leu His Thr Trp Gln Lys Cys Ser Asn Arg Leu Val 505 Asn Ser Gly Lys Val Lys Arg Glu Lys Arg Gly Asn Gly Trp Val Lys 520 Pro Gly Lys Gly Ser Glu Thr Val Val 530 535 <210> 50

<211> 537

<212> PRT

<213> Mouse

<400> 50

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 Phe
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 Ile
 Gln
 Tyr
 Gly
 Cys
 Ser
 Ser
 Gln
 Leu
 Gln
 Pro

 Thr
 Phe
 Thr
 Pro

Phe Leu Cys Ser Val Tyr Val Pro Met Cys Thr Glu Lys Ile Asn Ile Pro Ile Gly Pro Cys Gly Gly Met Cys Leu Ser Val Lys Arg Arg Cys Glu Pro Val Leu Arg Glu Phe Gly Phe Ala Trp Pro Asp Thr Leu Asn Cys Ser Lys Phe Pro Pro Gln Asn Asp His Asn His Met Cys Met Glu Gly Pro Gly Asp Glu Glu Val Pro Leu Pro His Lys Thr Pro Ile Gln Pro Gly Glu Glu Cys His Ser Val Gly Ser Asn Ser Asp Gln Tyr Ile Trp Val Lys Arg Ser Leu Asn Cys Val Leu Lys Cys Gly Tyr Asp Ala Gly Leu Tyr Ser Arg Ser Ala Lys Glu Phe Thr Asp Ile Trp Met Ala Val Trp Ala Ser Leu Cys Phe Ile Ser Thr Thr Phe Thr Val Leu Thr Phe Leu Ile Asp Ser Ser Arg Phe Ser Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Met Cys Tyr Asn Ile Tyr Ser Ile Ala Tyr Ile Val Arg Leu Thr Val Gly Arg Glu Arg Ile Ser Cys Asp Phe Glu Glu Ala Ala Glu Pro Val Leu Ile Gln Glu Gly Leu Lys Asn Thr Gly Cys Ala Ile Ile Phe Leu Leu Met Tyr Phe Phe Gly Met Ala Ser Ser Ile Trp Trp Val Ile Leu Thr Leu Thr Trp Phe Leu Ala Ala Gly Leu Lys Trp Gly His Glu Ala Ile Glu Met His Ser Ser Tyr Phe His Ile Ala Ala Trp Ala Ile Pro Ala Val Lys Thr Ile Val Ile Leu Ile Met Arg Leu Val Asp Ala Asp Glu Leu Thr Gly Leu Cys Tyr Val Gly Asn Gln Asn Leu Asp Ala Leu Thr Gly Phe Val Val Ala Pro Leu Phe Thr Tyr Leu Val Ile Gly Thr Leu Phe Ile Ala Ala Gly Leu Val Ala Leu Phe Lys Ile Arg Ser Asn Leu Gln Lys Asp Gly Thr Lys Thr Asp Lys Leu Glu Arg Leu Met Val Lys Ile Gly Val Phe Ser Val Leu Tyr Thr Val Pro Ala Thr Cys Val Ile Ala Cys Tyr Phe Tyr Glu Ile Ser Asn Trp Ala Leu Phe Arg Tyr Ser Ala Asp Asp Ser Asn Met Ala Val Glu Met Leu Lys Ile Phe Met Ser Leu Leu Val Gly Ile Thr Ser Gly Met Trp Ile Trp Ser Ala Lys Thr Leu His Thr Trp Gln Lys Cys Ser Asn Arg Leu Val Asn Ser Gly Lys Val Lys Arg Glu Lys Arg Gly Asn Gly Trp Val Lys Pro Gly Lys Gly Asn Glu Thr Val Val

<210> 51 <211> 585 <212> PRT <213> Homo sapiens

<400> 51

Met Ala Arg Pro Asp Pro Ser Ala Pro Pro Ser Leu Leu Leu Leu Leu -Leu Ala Gln Leu Val Gly Arg Ala Ala Ala Ser Lys Ala Pro Val 25 Cys Gln Glu Ile Thr Val Pro Met Cys Arg Gly Ile Gly Tyr Asn Leu 40 Thr His Met Pro Asn Gln Phe Asn His Asp Thr Gln Asp Glu Ala Gly 55 Leu Glu Val His Gln Phe Trp Pro Leu Val Glu Ile Gln Cys Ser Pro Asp Leu Arg Phe Phe Leu Cys Thr Met Tyr Thr Pro Ile Cys Leu Pro 85 90 Asp Tyr His Lys Pro Leu Pro Pro Cys Arg Ser Val Cys Glu Arg Ala 105 Lys Ala Gly Cys Ser Pro Leu Met Arg Gln Tyr Gly Phe Ala Trp Pro 120 Glu Arg Met Ser Cys Asp Arg Leu Pro Val Leu Gly Arg Asp Ala Glu 135 140 Val Leu Cys Met Asp Tyr Asn Arg Ser Glu Ala Thr Thr Ala Pro Pro 150 155 Arg Pro Phe Pro Ala Lys Pro Thr Leu Pro Gly Pro Pro Gly Ala Pro 165 170 Ala Ser Gly Gly Glu Cys Pro Ala Gly Gly Pro Phe Val Cys Lys Cys 185 Arg Glu Pro Phe Val Pro Ile Leu Lys Glu Ser His Pro Leu Tyr Asn 200 205 Lys Val Arg Thr Gly Gln Val Pro Asn Cys Ala Val Pro Cys Tyr Gln 215 220 Pro Ser Phe Ser Ala Asp Glu Arg Thr Phe Ala Thr Phe Trp Ile Gly 230 235 Leu Trp Ser Val Leu Cys Phe Ile Ser Thr Ser Thr Thr Val Ala Thr 250 245 Phe Leu Ile Asp Met Asp Thr Phe Arg Tyr Pro Glu Arg Pro Ile Ile 265 Phe Leu Ser Ala Cys Tyr Leu Cys Val Ser Leu Gly Phe Leu Val Arg 280 285 Leu Val Val Gly His Ala Ser Val Ala Cys Ser Arg Glu His Asn His Ile His Tyr Glu Thr Thr Gly Pro Ala Leu Cys Thr Ile Val Phe Leu 310 315 Leu Val Tyr Phe Phe Gly Met Ala Ser Ser Ile Trp Trp Val Ile Leu 325 330 Ser Leu Thr Trp Phe Leu Ala Ala Met Lys Trp Gly Asn Glu Ala 345 Ile Ala Gly Tyr Gly Gln Tyr Phe His Leu Ala Ala Trp Leu Ile Pro 360 365 Ser Val Lys Ser Ile Thr Ala Leu Ala Leu Ser Ser Val Asp Gly Asp 375 Pro Val Ala Gly Ile Cys Tyr Val Gly Asn Gln Asn Leu Asn Ser Leu 385 390 395

Arg Arg Phe Val Leu Gly Pro Leu Val Leu Tyr Leu Leu Val Gly Thr 410 Leu Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Ser Val 425 Ile Lys Gln Gly Gly Thr Lys Thr Asp Lys Leu Glu Lys Leu Met Ile 440 Arg Ile Gly Ile Phe Thr Leu Leu Tyr Thr Val Pro Ala Ser Ile Val 455 460 Val Ala Cys Tyr Leu Tyr Glu Gln His Tyr Arg Glu Ser Trp Glu Ala 470 475 Ala Leu Thr Cys Ala Cys Pro Gly His Asp Thr Gly Gln Pro Arg Ala 485 490 Lys Pro Glu Tyr Trp Val Leu Met Leu Lys Tyr Phe Met Cys Leu Val 505 500 Val Gly Ile Thr Ser Gly Val Trp Ile Trp Ser Gly Lys Thr Val Glu 520 Ser Trp Arg Arg Phe Thr Ser Arg Cys Cys Cys Arg Pro Arg Arg Gly 535 540 His Lys Ser Gly Gly Ala Met Ala Gly Asp Tyr Pro Glu Ala Ser 555 Ala Ala Leu Thr Gly Arg Thr Gly Pro Pro Gly Pro Ala Ala Thr Tyr 565 570 His Lys Gln Val Ser Leu Ser His Val 580

<210> 52 <211> 706

<212> PRT

<213> Homo sapiens

<400> 52

Met Glu Met Phe Thr Phe Leu Leu Thr Cys Ile Phe Leu Pro Leu Leu Arg Gly His Ser Leu Phe Thr Cys Glu Pro Ile Thr Val Pro Arg Cys 25 Met Lys Met Ala Tyr Asn Met Thr Phe Phe Pro Asn Leu Met Gly His 40 Tyr Asp Gln Ser Ile Ala Ala Val Glu Met Glu His Phe Leu Pro Leu 55 60 Ala Asn Leu Glu Cys Ser Pro Asn Ile Glu Thr Phe Leu Cys Lys Ala 70 75 Phe Val Pro Thr Cys Ile Glu Gln Ile His Val Val Pro Pro Cys Arg Lys Leu Cys Glu Lys Val Tyr Ser Asp Cys Lys Lys Leu Ile Asp Thr 105 Phe Gly Ile Arg Trp Pro Glu Glu Leu Glu Cys Asp Arg Leu Gln Tyr 120 Cys Asp Glu Thr Val Pro Val Thr Phe Asp Pro His Thr Glu Phe Leu 135 140 Gly Pro Gln Lys Lys Thr Glu Gln Val Gln Arg Asp Ile Gly Phe Trp 150 155 Cys Pro Arg His Leu Lys Thr Ser Gly Gly Gln Gly Tyr Lys Phe Leu 165 170 Gly Ile Asp Gln Cys Ala Pro Pro Cys Pro Asn Met Tyr Phe Lys Ser 185 Asp Glu Leu Glu Phe Ala Lys Ser Phe Ile Gly Thr Val Ser Ile Phe 195 200 205

Cys Leu Cys Ala Thr Leu Phe Thr Phe Leu Thr Phe Leu Ile Asp Val 215 Arg Arg Phe Arg Tyr Pro Glu Arg Pro Ile Ile Tyr Tyr Ser Val Cys 230 Tyr Ser Ile Val Ser Leu Met Tyr Phe Ile Gly Phe Leu Leu Gly Asp 245 250 Ser Thr Ala Cys Asn Lys Ala Asp Glu Lys Leu Glu Leu Gly Asp Thr 265 Val Val Leu Gly Ser Gln Asn Lys Ala Cys Thr Val Leu Phe Met Leu . 280 Leu Tyr Phe Phe Thr Met Ala Gly Thr Val Trp Trp Val Ile Leu Thr 295 300 Ile Thr Trp Phe Leu Ala Ala Gly Arg Lys Trp Ser Cys Glu Ala Ile 310 315 Glu Gln Lys Ala Val Trp Phe His Ala Val Ala Trp Gly Thr Pro Gly 325 330 Phe Leu Thr Val Met Leu Leu Ala Met Asn Lys Val Glu Gly Asp Asn 345 Ile Ser Gly Val Cys Phe Val Gly Leu Tyr Asp Leu Asp Ala Ser Arg 360 Tyr Phe Val Leu Leu Pro Leu Cys Leu Cys Val Phe Val Gly Leu Ser 375 380 Leu Leu Leu Ala Gly Ile Ile Ser Leu Asn His Val Arg Gln Val Ile 390 395 Gln His Asp Gly Arg Asn Gln Glu Lys Leu Lys Lys Phe Met Ile Arg 405 410 Ile Gly Val Phe Ser Gly Leu Tyr Leu Val Pro Leu Val Thr Leu Leu 425 Gly Cys Tyr Val Tyr Glu Gln Val Asn Arg Ile Thr Trp Glu Ile Thr 440 Trp Val Ser Asp His Cys Arg Gln Tyr His Ile Pro Cys Pro Tyr Gln 455 Ala Lys Ala Lys Ala Arg Pro Glu Leu Ala Leu Phe Met Ile Lys Tyr 470 475 Leu Met Thr Leu Ile Val Gly Ile Ser Ala Val Phe Trp Val Gly Ser 485 490 Lys Lys Thr Cys Thr Glu Trp Ala Gly Phe Phe Lys Arg Asn Arg Lys 505 Arg Asp Pro Ile Ser Glu Ser Arg Arg Val Leu Gln Glu Ser Cys Glu · 520 525 Phe Phe Leu Lys His Asn Ser Lys Val Lys His Lys Lys His Tyr 535 Lys Pro Ser Ser His Lys Leu Lys Val Ile Ser Lys Ser Met Gly Thr 550 555 Ser Thr Gly Ala Thr Ala Asn His Gly Thr Ser Ala Val Ala Ile Thr 570 Ser His Asp Tyr Leu Gly Gln Glu Thr Leu Thr Glu Ile Gln Thr Ser 585 Pro Glu Thr Ser Met Arg Glu Val Lys Ala Asp Gly Ala Ser Thr Pro 600 605 Arg Leu Arg Glu Gln Asp Cys Gly Glu Pro Ala Ser Pro Ala Ala Ser 615 620 Ile Ser Arg Leu Ser Gly Glu Gln Val Asp Gly Lys Gly Gln Ala Gly 630 635 Ser Val Ser Glu Ser Ala Arg Ser Glu Gly Arg Ile Ser Pro Lys Ser 650 Asp Ile Thr Asp Thr Gly Leu Ala Gln Ser Asn Asn Leu Gln Val Pro

660 665 Ser Ser Ser Glu Pro Ser Ser Leu Lys Gly Ser Thr Ser Leu Leu Val 680 His Pro Val Ser Gly Val Arg Lys Glu Gln Gly Gly Cys His Ser 690 695 700 Asp Thr 705 <210> 53 <211> 709 <212> PRT <213> Mouse <400> 53 Met Glu Arg Ser Pro Phe Leu Leu Ala Cys Ile Leu Leu Pro Leu Val Arg Gly His Ser Leu Phe Thr Cys Glu Pro Ile Thr Val Pro Arg Cys 25 20 Met Lys Met Thr Tyr Asn Met Thr Phe Phe Pro Asn Leu Met Gly His 40 Tyr Asp Gln Gly Ile Ala Ala Val Glu Met Gly His Phe Leu His Leu 55 60 Ala Asn Leu Glu Cys Ser Pro Asn Ile Glu Met Phe Leu Cys Gln Ala 70 75 Phe Ile Pro Thr Cys Thr Glu Gln Ile His Val Val Leu Pro Cys Arg 85 90 Lys Leu Cys Glu Lys Ile Val Ser Asp Cys Lys Leu Met Asp Thr 105 Phe Gly Ile Arg Trp Pro Glu Glu Leu Glu Cys Asn Arg Leu Pro His 120 125 Cys Asp Asp Thr Val Pro Val Thr Ser His Pro His Thr Glu Leu Ser 135 140 Gly Pro Gln Lys Lys Ser Asp Gln Val Pro Arg Asp Ile Gly Phe Trp 150 155 Cys Pro Lys His Leu Arg Thr Ser Gly Asp Gln Gly Tyr Arg Phe Leu 165 170 Gly Ile Glu Gln Cys Ala Pro Pro Cys Pro Asn Met Tyr Phe Lys Ser 180 185 190 Asp Glu Leu Asp Phe Ala Lys Ser Phe Ile Gly Ile Val Ser Ile Phe 200 205 Cys Leu Cys Ala Thr Leu Phe Thr Phe Leu Thr Phe Leu Ile Asp Val 215 220 Arg Arg Phe Arg Tyr Pro Glu Arg Pro Ile Ile Tyr Tyr Ser Val Cys 235 Tyr Ser Ile Val Ser Leu Met Tyr Phe Val Gly Phe Leu Leu Gly Asn 245 250 Ser Thr Ala Cys Asn Lys Ala Asp Glu Lys Leu Glu Leu Gly Asp Thr 265 Val Val Leu Gly Ser Lys Asn Lys Ala Cys Ser Val Val Phe Met Phe 280 285 Leu Tyr Phe Phe Thr Met Ala Gly Thr Val Trp Trp Val Ile Leu Thr 295 300 Ile Thr Trp Phe Leu Ala Ala Gly Arg Lys Trp Ser Cys Glu Ala Ile 310 315 Glu Gln Lys Ala Val Trp Phe His Ala Val Ala Trp Gly Ala Pro Gly 325 330 Phe Leu Thr Val Met Leu Leu Ala Met Asn Lys Val Glu Gly Asp Asn

340 345 Ile Ser Gly Val Cys Phe Val Gly Leu Tyr Asp Leu Asp Ala Ser Arg 360 Tyr Phe Val Leu Leu Pro Leu Cys Leu Cys Val Phe Val Gly Leu Ser 375 380 Leu Leu Leu Ala Gly Ile Ile Ser Leu Asn His Val Arg Gln Val Ile 390 395 Gln His Asp Gly Arg Asn Gln Glu Lys Leu Lys Lys Phe Met Ile Arg 410 Ile Gly Val Phe Ser Gly Leu Tyr Leu Val Pro Leu Val Thr Leu Leu 425 420 Gly Cys Tyr Val Tyr Glu Leu Val Asn Arg Ile Thr Trp Glu Met Thr 440 Trp Phe Ser Asp His Cys His Gln Tyr Arg Ile Pro Cys Pro Tyr Gln Ala Asn Pro Lys Ala Arg Pro Glu Leu Ala Leu Phe Met Ile Lys Tyr 470 475 Leu Met Thr Leu Ile Val Gly Ile Ser Ala Val Phe Trp Val Gly Ser Lys Lys Thr Cys Thr Glu Trp Ala Gly Phe Phe Lys Arg Asn Arg Lys 505 Arg Asp Pro Ile Ser Glu Ser Arg Arg Val Leu Gln Glu Ser Cys Glu 520 Phe Phe Leu Lys His Asn Ser Lys Val Lys His Lys Lys His Gly 535 540 Ala Pro Gly Pro His Arg Leu Lys Val Ile Ser Lys Ser Met Gly Thr 550 555 Ser Thr Gly Ala Thr Thr Asn His Gly Thr Ser Ala Met Ala Ile Ala 570 Asp His Asp Tyr Leu Gly Gln Glu Thr Ser Thr Glu Val His Thr Ser 580 585 Pro Glu Ala Ser Val Lys Glu Gly Arg Ala Asp Arg Ala Asn Thr Pro 600 Ser Ala Lys Asp Arg Asp Cys Gly Glu Ser Ala Gly Pro Ser Ser Lys 615 620 Leu Ser Gly Asn Arg Asn Gly Arg Glu Ser Arg Ala Gly Gly Leu Lys 630 635 Glu Arg Ser Asn Gly Ser Glu Gly Ala Pro Ser Glu Gly Arg Val Ser 645 650 Pro Lys Ser Ser Val Pro Glu Thr Gly Leu Ile Asp Cys Ser Thr Ser 665 Gln Ala Ala Ser Ser Pro Glu Pro Thr Ser Leu Lys Gly Ser Thr Ser 680 Leu Pro Val His Ser Ala Ser Arg Ala Arg Lys Glu Gln Gly Ala Gly 695 Ser His Ser Asp Ala 705 <210> 54 <211> 574 <212> PRT <213> Homo sapiens

<400> 54

Met Arg Asp Pro Gly Ala Ala Ala Pro Leu Ser Ser Leu Gly Leu Cys
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Ala Leu Val Leu Ala Leu Leu Gly Ala Leu Ser Ala Gly Ala Gly Ala

Gln Pro Tyr His Gly Glu Lys Gly Ile Ser Val Pro Asp His Gly Phe Cys Gln Pro Ile Ser Ile Pro Leu Cys Thr Asp Ile Ala Tyr Asn Gln Thr Ile Leu Pro Asn Leu Leu Gly His Thr Asn Gln Glu Asp Ala Gly Leu Glu Val His Gln Phe Tyr Pro Leu Val Lys Val Gln Cys Ser Pro Glu Leu Arg Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val Leu Asp Gln Ala Ile Pro Pro Cys Arg Ser Leu Cys Glu Arg Ala Arg Gln Gly Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln Trp Pro Glu Arg Leu Arg Cys Glu Asn Phe Pro Val His Gly Ala Gly Glu Ile Cys Val Gly Gln Asn Thr Ser Asp Gly Ser Gly Gly Pro Gly Gly Pro Thr Ala Tyr Pro Thr Ala Pro Tyr Leu Pro Asp Leu Pro Phe Thr Ala Leu Pro Pro Gly Ala Ser Asp Gly Arg Gly Arg Pro Ala Phe Pro Phe Ser Cys Pro Arg Gln Leu Lys Val Pro Pro Tyr Leu Gly Tyr Arg Phe Leu Gly Glu Arg Asp Cys Gly Ala Pro Cys Glu Pro Gly Arg Ala Asn Gly Leu Met Tyr Phe Lys Glu Glu Glu Arg Arg Phe Ala Arg Leu Trp Val Gly Val Trp Ser Val Leu Cys Cys Ala Ser Thr Leu Phe Thr Val Leu Thr Tyr Leu Val Asp Met Arg Arg Phe Ser Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Gly Cys Tyr Phe Met Val Ala Val Ala His Val Ala Gly Phe Leu Leu Glu Asp Arg Ala Val Cys Val Glu Arg Phe Ser Asp Asp Gly Tyr Arg Thr Val Ala Gln Gly Thr Lys Lys Glu Gly Cys Thr Ile Leu Phe Met Val Leu Tyr Phe Phe Gly Met Ala Ser Ser Ile Trp Trp Val Ile Leu Ser Leu Thr Trp Phe Leu Ala Ala Gly Met Lys Trp Gly His Glu Ala Ile Glu Ala Asn Ser Gln Tyr Phe His Leu Ala Ala Trp Ala Val Pro Ala Val Lys Thr Ile Thr Ile Leu Ala Met Gly Gln Val Asp Gly Asp Leu Leu Ser Gly Val Cys Tyr Val Gly Leu Ser Ser Val Asp Ala Leu Arg Gly Phe Val Leu Ala Pro Leu Phe Val Tyr Leu Phe Ile Gly Thr Ser Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Thr Ile Met Lys His Asp Gly Thr Lys Thr Glu Lys Leu Glu Lys Leu Met Val Arg Ile Gly Val Phe Ser Val Leu Tyr Thr Val

 Pro
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<213> Mouse

<400> 55

Met Arg Gly Pro Gly Thr Ala Ala Ser His Ser Pro Leu Gly Leu Cys Ala Leu Val Leu Ala Leu Leu Gly Ala Leu Pro Thr Asp Thr Arg Ala Gln Pro Tyr His Gly Glu Lys Gly Ile Ser Val Pro Asp His Gly Phe 40 Cys Gln Pro Ile Ser Ile Pro Leu Cys Thr Asp Ile Ala Tyr Asn Gln Thr Ile Leu Pro Asn Leu Leu Gly His Thr Asn Gln Glu Asp Ala Gly Leu Glu Val His Gln Phe Tyr Pro Leu Val Lys Val Gln Cys Ser Pro 90 Glu Leu Arg Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val 105 Leu Asp Gln Ala Ile Pro Pro Cys Arg Ser Leu Cys Glu Arg Ala Arg 120 Gln Gly Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln Trp Pro Glu 135 Arg Leu Arg Cys Glu Asn Phe Pro Val His Gly Ala Gly Glu Ile Cys 150 155 Val Gly Gln Asn Thr Ser Asp Gly Ser Gly Gly Ala Gly Gly Ser Pro 170 Thr Ala Tyr Pro Thr Ala Pro Tyr Leu Pro Asp Pro Pro Phe Thr Ala 185 Met Ser Pro Ser Asp Gly Arg Gly Arg Leu Ser Phe Pro Phe Ser Cys 200 Pro Arg Gln Leu Lys Val Pro Pro Tyr Leu Gly Tyr Arg Phe Leu Gly 215 220 Glu Arg Asp Cys Gly Ala Pro Cys Glu Pro Gly Arg Ala Asn Gly Leu 230 235 Met Tyr Phe Lys Glu Glu Glu Arg Arg Phe Ala Arg Leu Trp Val Gly 250 Val Trp Ser Val Leu Ser Cys Ala Ser Thr Leu Phe Thr Val Leu Thr 260 265 Tyr Leu Val Asp Met Arg Arg Phe Ser Tyr Pro Glu Arg Pro Ile Ile 280 Phe Leu Ser Gly Cys Tyr Phe Met Val Ala Val Ala His Val Ala Gly 290 295 300



Phe Leu Leu Glu Asp Arg Ala Val Cys Val Glu Arg Phe Ser Asp Asp 310 315 Gly Tyr Arg Thr Val Ala Gln Gly Thr Lys Lys Glu Gly Cys Thr Ile 325 330 Leu Phe Met Val Leu Tyr Phe Phe Gly Met Ala Ser Ser Ile Trp Trp 345 Val Ile Leu Ser Leu Thr Trp Phe Leu Ala Ala Gly Met Lys Trp Gly 360 His Glu Ala Ile Glu Ala Asn Ser Gln Tyr Phe His Leu Ala Ala Trp 375 380 Ala Val Pro Ala Val Lys Thr Ile Thr Ile Leu Ala Met Gly Gln Val 390 395 Asp Gly Asp Leu Leu Ser Gly Val Cys Tyr Val Gly Leu Ser Ser Val 410 Asp Ala Leu Arg Gly Phe Val Leu Ala Pro Leu Phe Val Tyr Leu Phe 425 430 Ile Gly Thr Ser Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile 440 Arg Thr Ile Met Lys His Asp Gly Thr Lys Thr Glu Lys Leu Glu Lys 455 460 Leu Met Val Arg Ile Gly Val Phe Ser Val Leu Tyr Thr Val Pro Ala 470 475 Thr Ile Val Leu Ala Cys Tyr Phe Tyr Glu Gln Ala Phe Arg Glu His 485 490 Trp Glu Arg Thr Trp Leu Leu Gln Thr Cys Lys Ser Tyr Ala Val Pro 500 505 Cys Pro Pro Arg His Phe Ser Pro Met Ser Pro Asp Phe Thr Val Phe 520 525 Met Ile Lys Tyr Leu Met Thr Met Ile Val Gly Ile Thr Thr Gly Phe 535 Trp Ile Trp Ser Gly Lys Thr Leu Gln Ser Trp Arg Arg Phe Tyr His 550 555 Arg Leu Ser His Ser Ser Lys Gly Glu Thr Ala Val 565

<210> 56

<211> 694

<212> PRT

<213> Homo sapiens

<400> 56

Met Glu Trp Gly Tyr Leu Leu Glu Val Thr Ser Leu Leu Ala Ala Leu Ala Leu Leu Gln Arg Ser Ser Gly Ala Ala Ala Ala Ser Ala Lys Glu 25 Leu Ala Cys Gln Glu Ile Thr Val Pro Leu Cys Lys Gly Ile Gly Tyr Asn Tyr Thr Tyr Met Pro Asn Gln Phe Asn His Asp Thr Gln Asp Glu 55 Ala Gly Leu Glu Val His Gln Phe Trp Pro Leu Val Glu Ile Gln Cys 75 Ser Pro Asp Leu Lys Phe Phe Leu Cys Ser Met Tyr Thr Pro Ile Cys 85 90 Leu Glu Asp Tyr Lys Lys Pro Leu Pro Pro Cys Arg Ser Val Cys Glu 105 Arg Ala Lys Ala Gly Cys Ala Pro Leu Met Arg Gln Tyr Gly Phe Ala 115 120 125

Trp Pro Asp Arg Met Arg Cys Asp Arg Leu Pro Glu Gln Gly Asn Pro 135 Asp Thr Leu Cys Met Asp Tyr Asn Arg Thr Asp Leu Thr Thr Ala Ala 150 155 Pro Ser Pro Pro Arg Arg Leu Pro Pro Pro Pro Gly Glu Gln Pro 170 Pro Ser Gly Ser Gly His Gly Arg Pro Pro Gly Ala Arg Pro Pro His 185 Arg Gly Gly Gly Gly Gly Gly Gly Asp Ala Ala Pro Pro 200 Ala Arg Gly Gly Gly Gly Gly Lys Ala Arg Pro Pro Gly Gly Gly 215 220 Ala Ala Pro Cys Glu Pro Gly Cys Gln Cys Arg Ala Pro Met Val Ser 230 235 Val Ser Ser Glu Arg His Pro Leu Tyr Asn Arg Val Lys Thr Gly Gln 245 250 Ile Ala Asn Cys Ala Leu Pro Cys His Asn Pro Phe Phe Ser Gln Asp 260 265 270 Glu Arg Ala Phe Thr Val Phe Trp Ile Gly Leu Trp Ser Val Leu Cys 280 Phe Val Ser Thr Phe Ala Thr Val Ser Thr Phe Leu Ile Asp Met Glu 295 300 Arg Phe Lys Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Ala Cys Tyr · 315 310 Leu Phe Val Ser Val Gly Tyr Leu Val Arg Leu Val Ala Gly His Glu 325 330 Lys Val Ala Cys Ser Gly Gly Ala Pro Gly Ala Gly Gly Ala Gly Gly 345 Ala Gly Gly Ala Ala Gly Ala Gly Ala Gly Ala Gly Ala Gly Gly Pro Gly Gly Arg Gly Glu Tyr Glu Glu Leu Gly Ala Val Glu Gln 375 His Val Arg Tyr Glu Thr Thr Gly Pro Ala Leu Cys Thr Val Val Phe 390 395 Leu Leu Val Tyr Phe Phe Gly Met Ala Ser Ser Ile Trp Trp Val Ile 405 410 Leu Ser Leu Thr Trp Phe Leu Ala Ala Gly Met Lys Trp Gly Asn Glu 420 425 430 Ala Ile Ala Gly Tyr Ser Gln Tyr Phe His Leu Ala Ala Trp Leu Val 440 Pro Ser Val Lys Ser Ile Ala Val Leu Ala Leu Ser Ser Val Asp Gly 455 460 Asp Pro Val Ala Gly Ile Cys Tyr Val Gly Asn Gln Ser Leu Asp Asn 470 475 Leu Arg Gly Phe Val Leu Ala Pro Leu Val Ile Tyr Leu Phe Ile Gly 485 490 Thr Met Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Ser 505 Val Ile Lys Gln Gln Asp Gly Pro Thr Lys Thr His Lys Leu Glu Lys 520 525 Leu Met Ile Arg Leu Gly Leu Phe Thr Val Leu Tyr Thr Val Pro Ala 535 540 Ala Val Val Ala Cys Leu Phe Tyr Glu Gln His Asn Arg Pro Arg 555 550 Trp Glu Ala Thr His Asn Cys Pro Cys Leu Arg Asp Leu Gln Pro Asp Gln Ala Arg Arg Pro Asp Tyr Ala Val Phe Met Leu Lys Tyr Phe Met

580 585 Cys Leu Val Val Gly Ile Thr Ser Gly Val Trp Val Trp Ser Gly Lys 600 Thr Leu Glu Ser Trp Arg Ser Leu Cys Thr Arg Cys Cys Trp Ala Ser 615 620 Lys Gly Ala Ala Val Gly Gly Gly Ala Gly Ala Thr Ala Ala Gly Gly 630 635 Gly Gly Pro Gly Gly Gly Gly Gly Gly Pro Gly Gly Gly Gly 645 650 Gly Pro Gly Gly Gly Gly Ser Leu Tyr Ser Asp Val Ser Thr Gly 665 Leu Thr Trp Arg Ser Gly Thr Ala Ser Ser Val Ser Tyr Pro Lys Gln 680 Met Pro Leu Ser Gln Val 690 <210> 57 <211> 685

<211> 665 <212> PRT <213> Mouse

<400> 57

Met Glu Trp Gly Tyr Leu Leu Glu Val Thr Ser Leu Leu Ala Ala Leu 10 Ala Val Leu Gln Arg Ser Ser Gly Ala Ala Ala Ser Ala Lys Glu 25 Leu Ala Cys Gln Glu Ile Thr Val Pro Leu Cys Lys Gly Ile Gly Tyr Asn Tyr Thr Tyr Met Pro Asn Gln Phe Asn His Asp Thr Gln Asp Glu Ala Gly Leu Glu Val His Gln Phe Trp Pro Leu Val Glu Ile Gln Cys 70 75 Ser Pro Asp Leu Lys Phe Phe Leu Cys Ser Met Tyr Thr Pro Ile Cys Leu Glu Asp Tyr Lys Lys Pro Leu Pro Pro Cys Arg Ser Val Cys Glu 105 Arg Ala Lys Ala Gly Cys Ala Pro Leu Met Arg Gln Tyr Gly Phe Ala 120 125 Trp Pro Asp Arg Met Arg Cys Asp Arg Leu Pro Glu Gln Gly Asn Pro 135 140 Asp Thr Leu Cys Met Asp Tyr Asn Arg Thr Asp Leu Thr Thr Ala Ala 155 150 Pro Ser Pro Pro Arg Arg Leu Pro Pro Pro Pro Pro Gly Glu Gln 170 Pro Pro Ser Gly Ser Gly His Ser Arg Pro Pro Gly Ala Arg Pro Pro 185 His Arg Gly Gly Ser Ser Arg Gly Ser Gly Asp Ala Ala Ala Pro 200 Pro Ser Arg Gly Gly Lys Ala Arg Pro Pro Gly Gly Gly Ala Ala Pro 215 220 Cys Glu Pro Gly Cys Gln Cys Arg Ala Pro Met Val Ser Val Ser Ser 230 235 Glu Arg His Pro Leu Tyr Asn Arg Val Lys Thr Gly Gln Ile Ala Asn 250 Cys Ala Leu Pro Cys His Asn Pro Phe Phe Ser Gln Asp Glu Arg Ala 265 Phe Thr Val Phe Trp Ile Gly Leu Trp Ser Val Leu Cys Phe Val Ser

280 285 275 Thr Phe Ala Thr Val Ser Thr Phe Leu Ile Asp Met Glu Arg Phe Lys 295 Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Ala Cys Tyr Leu Phe Val 310 315 Ser Val Gly Tyr Leu Val Arg Leu Val Ala Gly His Glu Lys Val Ala 325 330 Cys Ser Gly Gly Ala Pro Gly Ala Gly Gly Arg Gly Gly Ala Gly Gly 345 Ala Ala Ala Gly Ala Gly Ala Gly Arg Gly Ala Ser Ser Pro 360 365 Gly Ala Arg Gly Glu Tyr Glu Glu Leu Gly Ala Val Glu Gln His Val 375 380 Arg Tyr Glu Thr Thr Gly Pro Ala Leu Cys Thr Val Val Phe Leu Leu 390 395 Val Tyr Phe Phe Gly Met Ala Ser Ser Ile Trp Trp Val Ile Leu Ser 405 410 Leu Thr Trp Phe Leu Ala Ala Gly Met Lys Trp Gly Asn Glu Ala Ile 425 Ala Gly Tyr Ser Gln Tyr Phe His Leu Ala Ala Trp Leu Val Pro Ser 440 Val Lys Ser Ile Ala Val Leu Ala Leu Ser Ser Val Asp Gly Asp Pro 455 460 Val Ala Gly Ile Cys Tyr Val Gly Asn Gln Ser Leu Asp Asn Leu Arg 470 475 Gly Phe Val Leu Ala Pro Leu Val Ile Tyr Leu Phe Ile Gly Thr Met 485 490 Phe Leu Leu Ala Gly Phe Val Ser Leu Phe Arg Ile Arg Ser Val Ile 505 Lys Gln Gln Gly Gly Pro Thr Lys Thr His Lys Leu Glu Lys Leu Met 520 Ile Arg Leu Gly Leu Phe Thr Val Leu Tyr Thr Val Pro Ala Ala Val 535 Val Val Ala Cys Leu Phe Tyr Glu Gln His Asn Arg Pro Arg Trp Glu 550 555 Ala Thr His Asn Cys Pro Cys Leu Arq Asp Leu Gln Pro Asp Gln Ala 570 565 Arg Arg Pro Asp Tyr Ala Val Phe Met Leu Lys Tyr Phe Met Cys Leu 585 Val Val Gly Ile Thr Ser Gly Val Trp Val Trp Ser Gly Lys Thr Leu Glu Ser Trp Arg Ala Leu Cys Thr Arg Cys Cys Trp Ala Ser Lys Gly 615 Ala Ala Val Gly Ala Gly Ala Gly Gly Ser Gly Pro Gly Gly Ser Gly 635 Pro Gly Pro Gly Gly Gly Gly His Gly Gly Gly Gly Ser Leu 650 Tyr Ser Asp Val Ser Thr Gly Leu Thr Trp Arg Ser Gly Thr Ala Ser 665 Ser Val Ser Tyr Pro Lys Gln Met Pro Leu Ser Gln Val 680

<210> 58

<211> 591

<212> PRT

<213> Homo sapiens

<400> 58 Met Ala Val Ala Pro Leu Arg Gly Ala Leu Leu Leu Trp Gln Leu Leu Ala Ala Gly Gly Ala Ala Leu Glu Ile Gly Arg Phe Asp Pro Glu Arg 25 Gly Arg Gly Ala Ala Pro Cys Gln Ala Val Glu Ile Pro Met Cys Arg Gly Ile Gly Tyr Asn Leu Thr Arg Met Pro Asn Leu Leu Gly His Thr 55 Ser Gln Gly Glu Ala Ala Ala Glu Leu Ala Glu Phe Ala Pro Leu Val 75 Gln Tyr Gly Cys His Ser His Leu Arg Phe Phe Leu Cys Ser Leu Tyr 90 Ala Pro Met Cys Thr Asp Gln Val Ser Thr Pro Ile Pro Ala Cys Arg 105 Pro Met Cys Glu Gln Ala Arg Leu Arg Cys Ala Pro Ile Met Glu Gln Phe Asn Phe Gly Trp Pro Asp Ser Leu Asp Cys Ala Arg Leu Pro Thr 135 Arg Asn Asp Pro His Ala Leu Cys Met Glu Ala Pro Glu Asn Ala Thr 150 155 Ala Gly Pro Ala Glu Pro His Lys Gly Leu Gly Met Leu Pro Val Ala 165 170 Pro Arg Pro Ala Arg Pro Pro Gly Asp Leu Gly Pro Gly Ala Gly Gly 185 180 Ser Gly Thr Cys Glu Asn Pro Glu Lys Phe Gln Tyr Val Glu Lys Ser 200 205 Arg Ser Cys Ala Pro Arg Cys Gly Pro Gly Val Glu Val Phe Trp Ser 215 220 Arg Arg Asp Lys Asp Phe Ala Leu Val Trp Met Ala Val Trp Ser Ala 230 235 Leu Cys Phe Phe Ser Thr Ala Phe Thr Val Leu Thr Phe Leu Leu Glu 250 245 Pro His Arg Phe Gln Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Met 265 Cys Tyr Asn Val Tyr Ser Leu Ala Phe Leu Ile Arg Ala Val Ala Gly 280 Ala Gln Ser Val Ala Cys Asp Gln Glu Ala Gly Ala Leu Tyr Val Ile 295 300 Gln Glu Gly Leu Glu Asn Thr Gly Cys Thr Leu Val Phe Leu Leu 310 315 Tyr Tyr Phe Gly Met Ala Ser Ser Leu Trp Trp Val Val Leu Thr Leu 325 330 Thr Trp Phe Leu Ala Ala Gly Lys Lys Trp Gly His Glu Ala Ile Glu Ala His Gly Ser Tyr Phe His Met Ala Ala Trp Gly Leu Pro Ala Leu 360 Lys Thr Ile Val Ile Leu Thr Leu Arg Lys Val Ala Gly Asp Glu Leu 375 Thr Gly Leu Cys Tyr Val Ala Ser Thr Asp Ala Ala Ala Leu Thr Gly 390 395 Phe Val Leu Val Pro Leu Ser Gly Tyr Leu Val Leu Gly Ser Ser Phe 405 410 Leu Leu Thr Gly Phe Val Ala Leu Phe His Ile Arg Lys Ile Met Lys 425 Thr Gly Gly Thr Asn Thr Glu Lys Leu Glu Lys Leu Met Val Lys Ile 435 440



Gly Val Phe Ser Ile Leu Tyr Thr Val Pro Ala Thr Cys Val Ile Val 455 Cys Tyr Val Tyr Glu Arg Leu Asn Met Asp Phe Trp Arg Leu Arg Ala 470 475 Thr Glu Gln Pro Cys Ala Ala Ala Ala Gly Pro Gly Gly Arg Arg Asp 485 490 Cys Ser Leu Pro Gly Gly Ser Val Pro Thr Val Ala Val Phe Met Leu 505 Lys Ile Phe Met Ser Leu Val Val Gly Ile Thr Ser Gly Val Trp Val 520 Trp Ser Ser Lys Thr Phe Gln Thr Trp Gln Ser Leu Cys Tyr Arg Lys 535 540 Ile Ala Ala Gly Arg Ala Arg Ala Lys Ala Cys Arg Ala Pro Gly Ser 550 . 555 Tyr Gly Arg Gly Thr His Cys His Tyr Lys Ala Pro Thr Val Val Leu 570 His Met Thr Lys Thr Asp Pro Ser Leu Glu Asn Pro Thr His Leu 580 585

<210> 59

<211> 591

<212> PRT

<213> Mouse

<400> 59

Met Ala Val Pro Pro Leu Leu Arg Gly Ala Leu Leu Trp Gln Leu Leu Ala Thr Gly Gly Ala Ala Leu Glu Ile Gly Arg Phe Asp Pro Glu Arg Gly Arg Gly Pro Ala Pro Cys Gln Ala Met Glu Ile Pro Met Cys 40 Arg Gly Ile Gly Tyr Asn Leu Thr Arg Met Pro Asn Leu Leu Gly His 55 60 Thr Ser Gln Gly Glu Ala Ala Gln Leu Ala Glu Phe Ser Pro Leu 75 Val Gln Tyr Gly Cys His Ser His Leu Arg Phe Phe Leu Cys Ser Leu 90 Tyr Ala Pro Met Cys Thr Asp Gln Val Ser Thr Pro Ile Pro Ala Cys 105 Arg Pro Met Cys Glu Gln Ala Arg Leu Arg Cys Ala Pro Ile Met Glu 120 Gln Phe Asn Phe Gly Trp Pro Asp Ser Leu Asp Cys Ala Arg Leu Pro 135 140 Thr Arg Asn Asp Pro His Ala Leu Cys Met Glu Ala Pro Glu Asn Thr 155 Ala Gly Pro Thr Glu Pro His Lys Gly Leu Gly Met Leu Pro Val Ala 165 170 Pro Arg Pro Ala Arg Pro Pro Gly Asp Ser Ala Pro Gly Pro Gly Ser 185 Gly Gly Thr Cys Asp Asn Pro Glu Lys Phe Gln Tyr Val Glu Lys Ser 200 Arg Ser Cys Ala Pro Arg Cys Gly Pro Gly Val Glu Val Phe Trp Ser 215 220 Arg Arg Asp Lys Asp Phe Ala Leu Val Trp Met Ala Val Trp Ser Ala 230 235 Leu Cys Phe Phe Ser Thr Ala Phe Thr Val Phe Thr Phe Leu Leu Glu 245 250

Pro His Arg Phe Gln Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Met 260 265 Cys Tyr Asn Val Tyr Ser Leu Ala Phe Leu Ile Arg Ala Val Ala Gly 280 Ala Gln Ser Val Ala Cys Asp Gln Glu Ala Gly Ala Leu Tyr Val Ile 295 Gln Glu Gly Leu Glu Asn Thr Gly Cys Thr Leu Val Phe Leu Leu Leu 310 315 Tyr Tyr Phe Gly Met Ala Ser Ser Leu Trp Trp Val Val Leu Thr Leu 325 330 Thr Trp Phe Leu Ala Ala Gly Lys Lys Trp Gly His Glu Ala Ile Glu 340 345 Ala His Gly Ser Tyr Phe His Met Ala Ala Trp Gly Leu Pro Ala Leu 360 Lys Thr Ile Val Val Leu Thr Leu Arg Lys Val Ala Gly Asp Glu Leu 375 Thr Gly Leu Cys Tyr Val Ala Ser Met Asp Pro Ala Ala Leu Thr Gly 390 395 Phe Val Leu Val Pro Leu Ser Cys Tyr Leu Val Leu Gly Thr Ser Phe 410 Leu Leu Thr Gly Phe Val Ala Leu Phe His Ile Arg Lys Ile Met Lys 425 420 Thr Gly Gly Thr Asn Thr Glu Lys Leu Glu Lys Leu Met Val Lys Ile 435 440 445 Gly Val Phe Ser Ile Leu Tyr Thr Val Pro Ala Thr Cys Val Ile Val 455 460 Cys Tyr Val Tyr Glu Arg Leu Asn Met Asp Phe Trp Arg Leu Arg Ala 470 475 Thr Glu Gln Pro Cys Thr Ala Ala Thr Val Pro Gly Gly Arg Arg Asp 490 Cys Ser Leu Pro Gly Gly Ser Val Pro Thr Val Ala Val Phe Met Leu 505 Lys Ile Phe Met Ser Leu Val Val Gly Ile Thr Ser Gly Val Trp Val 520 Trp Ser Ser Lys Thr Phe Gln Thr Trp Gln Ser Leu Cys Tyr Arg Lys 535 540 Met Ala Ala Gly Arg Ala Arg Ala Lys Ala Cys Arg Thr Pro Gly Gly 550 555 Tyr Gly Arg Gly Thr His Cys His Tyr Lys Ala Pro Thr Val Val Leu 565 . 570 His Met Thr Lys Thr Asp Pro Ser Leu Glu Asn Pro Thr His Leu 585

<210> 60

<211> 581

<212> PRT

<213> Homo sapiens

<400> 60

 Met Gln Arg
 Pro Gly
 Pro Arg
 Leu
 Trp
 Leu
 Val
 Leu
 Gln
 Val
 Met
 Gly

 1
 5
 Leu
 Trp
 Leu
 Val
 Leu
 Gln
 Val
 Met
 Gly

 Ser
 Cys
 Ala
 Ala
 Ile
 Ser
 Ser
 Met
 Asp
 Met
 Glu
 Arg
 Pro
 Gly
 Asp
 Gly
 Asp
 Ile
 Gly
 Tyr
 Asn
 Asn
 45
 Asp
 Met
 Thr
 Arg
 Met
 Pro
 Asn
 Leu
 Met
 Gly
 His
 Glu
 Asn
 Glu
 Ala

 50
 55
 Trans
 From Arg
 From Arg</t

Ala Ile Gln Leu His Glu Phe Ala Pro Leu Val Glu Tyr Gly Cys His Gly His Leu Arg Phe Phe Leu Cys Ser Leu Tyr Ala Pro Met Cys Thr Glu Gln Val Ser Thr Pro Ile Pro Ala Cys Arg Val Met Cys Glu Gln 105 Ala Arg Leu Lys Cys Ser Pro Ile Met Glu Gln Phe Asn Phe Lys Trp 120 Pro Asp Ser Leu Asp Cys Arg Lys Leu Pro Asn Lys Asn Asp Pro Asn 135 Tyr Leu Cys Met Glu Ala Pro Asn Asn Gly Ser Asp Glu Pro Thr Arg 150 155 Gly Ser Gly Leu Phe Pro Pro Leu Phe Arg Pro Gln Arg Pro His Ser 165 170 Ala Gln Glu His Pro Leu Lys Asp Gly Gly Pro Gly Arg Gly Gly Cys 185 Asp Asn Pro Gly Lys Phe His His Val Glu Lys Ser Ala Ser Cys Ala 200 Pro Leu Cys Thr Pro Gly Val Asp Val Tyr Trp Ser Arg Glu Asp Lys 215 Arg Phe Ala Val Val Trp Leu Ala Ile Trp Ala Val Leu Cys Phe Phe 230 235 Ser Ser Ala Phe Thr Val Leu Thr Phe Leu Ile Asp Pro Ala Arg Phe 245 250 Arg Tyr Pro Glu Arg Pro Ile Ile Phe Leu Ser Met Cys Tyr Cys Val 260 265 Tyr Ser Val Gly Tyr Leu Ile Arg Leu Phe Ala Gly Ala Glu Ser Ile 280 Ala Cys Asp Arg Asp Ser Gly Gln Leu Tyr Val Ile Gln Glu Gly Leu 295 Glu Ser Thr Gly Cys Thr Leu Val Phe Leu Val Leu Tyr Tyr Phe Gly 310 315 Met Ala Ser Ser Leu Trp Trp Val Val Leu Thr Leu Thr Trp Phe Leu 330 Ala Ala Gly Lys Lys Trp Gly His Glu Ala Ile Glu Ala Asn Ser Ser 345 Tyr Phe His Leu Ala Ala Trp Ala Ile Pro Ala Val Lys Thr Ile Leu 360 Ile Leu Val Met Arg Arg Val Ala Gly Asp Glu Leu Thr Gly Val Cys 375 380 Tyr Val Gly Ser Met Asp Val Asn Ala Leu Thr Gly Phe Val Leu Ile 390 395 Pro Leu Ala Cys Tyr Leu Val Ile Gly Thr Ser Phe Ile Leu Ser Gly 410 Phe Val Ala Leu Phe His Ile Arg Arg Val Met Lys Thr Gly Glu 425 430 Asn Thr Asp Lys Leu Glu Lys Leu Met Val Arg Ile Gly Leu Phe Ser 440 Val Leu Tyr Thr Val Pro Ala Thr Cys Val Ile Ala Cys Tyr Phe Xaa 455 460 Glu His Leu Asn Met Asp Tyr Trp Lys Ile Leu Ala Ala Gln His Lys 470 475 Cys Lys Met Asn Asn Gln Thr Lys Thr Leu Asp Cys Leu Met Ala Ala 485 490 Ser Ile Pro Ala Val Glu Ile Phe Met Val Lys Ile Phe Met Leu Leu 505 Val Val Gly Ile Thr Ser Gly Met Trp Ile Trp Thr Ser Lys Thr Leu

520 Gln Ser Trp Gln Gln Val Cys Ser Arg Arg Leu Lys Lys Lys Ser Arg 535 Arg Lys Pro Ala Ser Val Ile Thr Ser Gly Gly Ile Tyr Lys Lys Ala 550 555 Gln His Pro Gln Lys Thr His His Gly Lys Tyr Glu Ile Pro Ala Gln 570 Ser Pro Thr Cys Val 580 <210> 61 <211> 319 <212> PRT <213> Homo sapiens <400> 61 Met Ala Glu Glu Ala Pro Lys Lys Ser Arg Ala Ala Gly Gly Gly 10 Ala Ser Trp Glu Leu Cys Ala Gly Ala Leu Ser Ala Arg Leu Ala Glu 25 Glu Gly Ser Gly Asp Ala Gly Gly Arg Arg Pro Pro Val Asp Pro 40 Arg Arg Leu Ala Arg Gln Leu Leu Leu Leu Trp Leu Leu Glu Ala Pro Leu Leu Gly Val Arg Ala Gln Ala Gly Gln Gly Pro Gly 70 75 . Gln Gly Pro Gly Pro Gly Gln Gln Pro Pro Pro Pro Gln Gln Gln Gln Ser Gly Gln Gln Tyr Asn Gly Glu Arg Gly Ile Ser Val Pro Asp 105 His Gly Tyr Cys Gln Pro Ile Ser Ile Pro Leu Cys Thr Asp Ile Ala 115 120 125 Tyr Asn Gln Thr Ile Met Pro Asn Leu Leu Gly His Thr Asn Gln Glu 135 Asp Ala Gly Leu Glu Val His Gln Phe Tyr Pro Leu Val Lys Val Gln 150 155 Cys Ser Ala Glu Leu Lys Phe Phe Leu Cys Ser Met Tyr Ala Pro Val 165 170 Cys Thr Val Leu Glu Gln Ala Leu Pro Pro Cys Arg Ser Leu Cys Glu 185 Arg Ala Arg Gln Gly Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln 200 Trp Pro Asp Thr Leu Lys Cys Glu Lys Phe Pro Val His Gly Ala Gly 220 215 Glu Leu Cys Val Gly Gln Asn Thr Ser Asp Lys Gly Thr Pro Thr Pro 230 235 Ser Leu Leu Pro Glu Phe Trp Thr Ser Asn Pro Gln His Gly Gly Gly 245 250 Gly His Arg Gly Gly Phe Pro Gly Gly Ala Gly Ala Ser Glu Arg Gly 265 Lys Phe Ser Cys Pro Arg Ala Leu Lys Val Pro Ser Tyr Leu Asn Tyr 280 His Phe Leu Gly Glu Lys Asp Cys Gly Ala Pro Cys Glu Pro Thr Lys 295 300 Val Tyr Gly Leu Met Tyr Phe Gly Pro Glu Glu Leu Arg Phe Ser

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<210> 62
<211> 314
<212> PRT
<213> Mouse
<400> 62
Met Ala Glu Glu Ala Ala Pro Ser Glu Ser Arg Ala Ala Gly Arg Leu
Ser Leu Glu Leu Cys Ala Glu Ala Leu Pro Gly Arg Arg Glu Glu Val
Gly His Glu Asp Thr Ala Ser His Arg Arg Pro Arg Ala Asp Pro Arg
                           40
Arg Trp Ala Ser Gly Leu Leu Leu Leu Trp Leu Leu Glu Ala Pro
                        55
Leu Leu Gly Val Arg Ala Gln Ala Gly Gln Val Ser Gly Pro
Gly Gln Gln Ala Pro Pro Pro Gln Pro Gln Gln Ser Gly Gln Gln
               85
                                   90
Tyr Asn Gly Glu Arg Gly Ile Ser Ile Pro Asp His Gly Tyr Cys Gln
                               105
Pro Ile Ser Ile Pro Leu Cys Thr Asp Met Ala Tyr Asn Gln Thr Ile
                           120
       115
Met Pro Asn Leu Leu Gly His Thr Asn Gln Glu Asp Ala Gly Leu Glu
                       135
                                           140
Val His Gln Phe Tyr Pro Leu Val Lys Val Gln Cys Ser Ala Glu Leu
                   150
                                       155
Lys Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val Leu Glu
               165
                                   170
Gln Ala Leu Pro Pro Cys Arg Ser Leu Cys Glu Arg Ala Arg Gln Gly
Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln Trp Pro Asp Thr Leu
                            200
Lys Cys Glu Lys Phe Pro Val His Gly Ala Gly Glu Leu Cys Val Gly
                       215
Gln Asn Thr Ser Asp Lys Gly Thr Pro Thr Pro Ser Leu Leu Pro Glu
                   230
                                       235
Phe Trp Thr Ser Asn Gly Gln His Gly Gly Gly Tyr Arg Gly Gly
               245
                                   250
Tyr Pro Gly Gly Ala Gly Thr Val Glu Arg Gly Lys Phe Ser Cys Pro-
           260
                               265
Arg Ala Leu Arg Val Pro Ser Tyr Leu Asn Tyr His Phe Leu Gly Glu
                           280
Lys Asp Cys Gly Ala Pro Cys Glu Pro Thr Lys Val Tyr Gly Leu Met
                        295
Tyr Phe Gly Pro Glu Glu Leu Arg Phe Ser
                    310
<210> 63
<211> 244
<212> PRT
<213> Homo sapiens
<400> 63
Met Arg Pro Arg Ser Ala Leu Pro Arg Leu Leu Pro Leu Leu Leu
Leu Pro Ala Ala Gly Pro Ala Gln Phe His Gly Glu Lys Gly Ile Ser
           20
                                25
```

Ile Pro Asp His Gly Phe Cys Gln Pro Ile Ser Ile Pro Leu Cys Thr 40 Asp Ile Ala Tyr Asn Gln Thr Ile Met Pro Asn Leu Leu Gly His Thr Asn Gln Glu Asp Ala Gly Leu Glu Val His Gln Phe Tyr Pro Leu Val 75 Lys Val Gln Cys Ser Pro Glu Leu Arg Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val Leu Glu Gln Ala Ile Pro Pro Cys Arg Ser 105 Ile Cys Glu Arg Ala Arg Gln Gly Cys Glu Ala Leu Met Asn Lys Phe 120 125 Gly Phe Gln Trp Pro Glu Arg Leu Arg Cys Glu His Phe Pro Arg His 135 Gly Ala Glu Gln Ile Cys Val Gly Gln Asn His Ser Glu Asp Gly Ala Pro Ala Leu Leu Thr Thr Ala Pro Pro Pro Gly Leu Gln Pro Gly Ala 170 165 Gly Gly Thr Pro Gly Gly Pro Gly Gly Gly Gly Ala Pro Pro Arg Tyr 185 Ala Thr Leu Glu His Pro Phe His Cys Pro Arg Val Leu Lys Val Pro 195 200 Ser Tyr Leu Ser Tyr Lys Phe Leu Gly Glu Arg Asp Cys Ala Ala Pro 215 220 Cys Glu Pro Ala Arg Pro Asp Gly Ser Met Phe Phe Ser Gln Glu Glu 230 235 Thr Arg Phe Ala

<210> 64

<211> 202

<212> PRT

<213> Homo sapiens

<400> 64

Met Ala Met Thr Trp Ile Val Phe Ser Leu Trp Pro Leu Thr Val Phe 10 Met Gly His Ile Gly Gly His Ser Leu Phe Ser Cys Glu Pro Ile Thr 25 Leu Arg Met Cys Gln Asp Leu Pro Tyr Asn Thr Thr Phe Met Pro Asn Leu Leu Asn His Tyr Asp Gln Gln Thr Ala Ala Leu Ala Met Glu Pro 55 Phe His Pro Met Val Asn Leu Asp Cys Ser Arg Asp Phe Arg Pro Phe Leu Cys Ala Leu Tyr Ala Pro Ile Cys Met Glu Tyr Gly Arg Val Thr 90 Leu Pro Cys Arg Arg Leu Cys Gln Arg Ala Tyr Ser Glu Cys Ser Lys 105 Leu Met Glu Met Phe Gly Val Pro Trp Pro Glu Asp Met Glu Cys Ser 120 Arg Phe Pro Asp Cys Asp Glu Pro Tyr Pro Arg Leu Val Asp Leu Asn 135 140 Leu Ala Gly Glu Pro Thr Glu Gly Ala Pro Val Ala Val Gln Arg Asp 155 Tyr Gly Phe Trp Cys Pro Arg Glu Leu Lys Ile Asp Pro Asp Leu Gly 165 170

Tyr Ser Phe Leu His Val Arg Asp Cys Ser Pro Pro Cys Pro Asn Met

180

Tyr Phe Arg Arg Glu Glu Leu Ser Phe Ala

195

200

210> 65

2211> 202

2212> PRT

2213> Mouse

400> 65

Met Ala Val Ser Trp Ile Val Phe Asp Leu Trp Leu Leu Thr Val Phe

Met Ala Val Ser Trp Ile Val Phe Asp Leu Trp Leu Leu Thr Val Phe . 10 Leu Gly Gln Ile Gly Gly His Ser Leu Phe Ser Cys Glu Pro Ile Thr Leu Arg Met Cys Gln Asp Leu Pro Tyr Asn Thr Thr Phe Met Pro Asn Leu Leu Asn His Tyr Asp Gln Gln Thr Ala Ala Leu Ala Met Glu Pro 55 Phe His Pro Met Val Asn Leu Asp Cys Ser Arg Asp Phe Arg Pro Phe Leu Cys Ala Leu Tyr Ala Pro Ile Cys Met Glu Tyr Gly Arg Val Thr Leu Pro Cys Arg Arg Leu Cys Gln Arg Ala Tyr Ser Glu Cys Ser Lys 105 Leu Met Glu Met Phe Gly Val Pro Trp Pro Glu Asp Met Glu Cys Ser 120 Arg Phe Pro Asp Cys Asp Glu Pro Tyr Pro Arg Leu Val Asp Leu Asn 135 140 Leu Val Gly Asp Pro Thr Glu Gly Ala Pro Val Ala Val Gln Arg Asp 150 155 Tyr Gly Phe Trp Cys Pro Arg Glu Leu Lys Ile Asp Pro Asp Leu Gly 165 170 Tyr Ser Phe Leu His Val Arg Asp Cys Ser Pro Pro Cys Pro Asn Met 185 Tyr Phe Arg Arg Glu Glu Leu Ser Phe Ala 195 200

<210> 66 <211> 219 <212> PRT <213> Homo sapiens

<400> 66

 Met Ala Trp Arg Gly Arg
 Gly Ala Gly Pro Ser Val Pro Gly Ala Pro Gly Gly 1
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 10
 15
 15

 Val Gly Leu Ser Leu Gly Leu Leu Leu Gln Leu Leu Leu Leu Leu Gly 20
 25
 30
 16
 15
 15

 Pro Ala Arg Gly Phe Gly Asp Glu Glu Glu Glu Arg Arg Cys Asp Pro Ile 35
 40
 45
 45
 45

 Arg Ile Ser Met Cys Gln Asn Leu Gly Tyr Asn Val Thr Lys Met Pro 50
 55
 60
 60

 Asn Leu Val Gly His Glu Leu Gln Thr Asp Ala Glu Leu Gln Leu Thr 65
 70
 75
 80

 Thr Phe Thr Pro Leu Ile Gln Tyr Gly Cys Ser Ser Gln Leu Gln Phe 85
 90
 95

 Phe Leu Cys Ser Val Tyr Val Pro Met Cys Thr Glu Lys Ile Asn Ile

```
105
Pro Ile Gly Pro Cys Gly Gly Met Cys Leu Ser Val Lys Arg Arg Cys
                            120
                                                125
Glu Pro Val Leu Lys Glu Phe Gly Phe Ala Trp Pro Glu Ser Leu Asn
                                           140
                        135
Cys Ser Lys Phe Pro Pro Gln Asn Asp His Asn His Met Cys Met Glu
                   150
                                       155
Gly Pro Gly Asp Glu Glu Val Pro Leu Pro His Lys Thr Pro Ile Gln
                165
                                   170
Pro Gly Glu Glu Cys His Ser Val Gly Thr Asn Ser Asp Gln Tyr Ile
                               185
Trp Val Lys Arg Ser Leu Asn Cys Val Leu Lys Cys Gly Tyr Asp Ala
                            200
Gly Leu Tyr Ser Arg Ser Ala Lys Glu Phe Thr
                        215
<210> 67
<211> 219
<212> PRT
<213> Mouse
<400> 67
Met Ala Trp Pro Gly Thr Gly Pro Ser Ser Arg Gly Ala Pro Gly Gly
Val Gly Leu Arg Leu Gly Leu Leu Gln Phe Leu Leu Leu Arg
                                25
Pro Thr Leu Gly Phe Gly Asp Glu Glu Arg Arg Cys Asp Pro Ile
Arg Ile Ala Met Cys Gln Asn Leu Gly Tyr Asn Val Thr Lys Met Pro
Asn Leu Val Gly His Glu Leu Gln Thr Asp Ala Glu Leu Gln Leu Thr
                    70
                                        75
Thr Phe Thr Pro Leu Ile Gln Tyr Gly Cys Ser Ser Gln Leu Gln Phe
                                    90
Phe Leu Cys Ser Val Tyr Val Pro Met Cys Thr Glu Lys Ile Asn Ile
                                105
Pro Ile Gly Pro Cys Gly Gly Met Cys Leu Ser Val Lys Arg Arg Cys
                            120
                                                125
Glu Pro Val Leu Arg Glu Phe Gly Phe Ala Trp Pro Asp Thr Leu Asn
                       135
                                           140
Cys Ser Lys Phe Pro Pro Gln Asn Asp His Asn His Met Cys Met Glu
                    150
                                        155
Gly Pro Gly Asp Glu Glu Val Pro Leu Pro His Lys Thr Pro Ile Gln
                                    170
Pro Gly Glu Glu Cys His Ser Val Gly Ser Asn Ser Asp Gln Tyr Ile
                               185
Trp Val Lys Arg Ser Leu Asn Cys Val Leu Lys Cys Gly Tyr Asp Ala
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Gly Leu Tyr Ser Arg Ser Ala Lys Glu Phe Thr
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                        215
<210> 68
<211> 235
<212> PRT
<213> Homo sapiens
<400> 68
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50

Met Ala Arg Pro Asp Pro Ser Ala Pro Pro Ser Leu Leu Leu Leu Leu Leu Ala Gln Leu Val Gly Arg Ala Ala Ala Ala Ser Lys Ala Pro Val Cys Gln Glu Ile Thr Val Pro Met Cys Arg Gly Ile Gly Tyr Asn Leu Thr His Met Pro Asn Gln Phe Asn His Asp Thr Gln Asp Glu Ala Gly 55 Leu Glu Val His Gln Phe Trp Pro Leu Val Glu Ile Gln Cys Ser Pro Asp Leu Arg Phe Phe Leu Cys Thr Met Tyr Thr Pro Ile Cys Leu Pro 85 90 . Asp Tyr His Lys Pro Leu Pro Pro Cys Arg Ser Val Cys Glu Arg Ala 105 Lys Ala Gly Cys Ser Pro Leu Met Arg Gln Tyr Gly Phe Ala Trp Pro 120 Glu Arg Met Ser Cys Asp Arg Leu Pro Val Leu Gly Arg Asp Ala Glu 135 140 Val Leu Cys Met Asp Tyr Asn Arg Ser Glu Ala Thr Thr Ala Pro Pro 150 155 Arg Pro Phe Pro Ala Lys Pro Thr Leu Pro Gly Pro Pro Gly Ala Pro 170 165 Ala Ser Gly Gly Glu Cys Pro Ala Gly Gly Pro Phe Val Cys Lys Cys 185 Arg Glu Pro Phe Val Pro Ile Leu Lys Glu Ser His Pro Leu Tyr Asn 200 205 Lys Val Arg Thr Gly Gln Val Pro Asn Cys Ala Val Pro Cys Tyr Gln 215 Pro Ser Phe Ser Ala Asp Glu Arg Thr Phe Ala

<210> 69

<211> 198

<212> PRT

<213> Homo sapiens

<400> 69

Met Glu Met Phe Thr Phe Leu Leu Thr Cys Ile Phe Leu Pro Leu Leu 10 Arg Gly His Ser Leu Phe Thr Cys Glu Pro Ile Thr Val Pro Arg Cys Met Lys Met Ala Tyr Asn Met Thr Phe Phe Pro Asn Leu Met Gly His Tyr Asp Gln Ser Ile Ala Ala Val Glu Met Glu His Phe Leu Pro Leu 55 Ala Asn Leu Glu Cys Ser Pro Asn Ile Glu Thr Phe Leu Cys Lys Ala 70 - 75 Phe Val Pro Thr Cys Ile Glu Gln Ile His Val Val Pro Pro Cys Arg Lys Leu Cys Glu Lys Val Tyr Ser Asp Cys Lys Lys Leu Ile Asp Thr 105 Phe Gly Ile Arg Trp Pro Glu Glu Leu Glu Cys Asp Arg Leu Gln Tyr 120 125 Cys Asp Glu Thr Val Pro Val Thr Phe Asp Pro His Thr Glu Phe Leu 135 140 Gly Pro Gln Lys Lys Thr Glu Gln Val Gln Arg Asp Ile Gly Phe Trp 145 150 155

Cys Pro Arg His Leu Lys Thr Ser Gly Gly Gln Gly Tyr Lys Phe Leu
165 170 175

Gly Ile Asp Gln Cys Ala Pro Pro Cys Pro Asn Met Tyr Phe Lys Ser
180 185 190

Asp Glu Leu Glu Phe Ala
195

<210> 70
<211> 198
<212> PRT
<213> Mouse

<400> 70

Met Glu Arg Ser Pro Phe Leu Leu Ala Cys Ile Leu Leu Pro Leu Val

Met Glu Arg Ser Pro Phe Leu Leu Ala Cys Ile Leu Leu Pro Leu Val Arg Gly His Ser Leu Phe Thr Cys Glu Pro Ile Thr Val Pro Arg Cys Met Lys Met Thr Tyr Asn Met Thr Phe Phe Pro Asn Leu Met Gly His 40 Tyr Asp Gln Gly Ile Ala Ala Val Glu Met Gly His Phe Leu His Leu 55 Ala Asn Leu Glu Cys Ser Pro Asn Ile Glu Met Phe Leu Cys Gln Ala Phe Ile Pro Thr Cys Thr Glu Gln Ile His Val Val Leu Pro Cys Arg 85 90 Lys Leu Cys Glu Lys Ile Val Ser Asp Cys Lys Lys Leu Met Asp Thr 105 Phe Gly Ile Arg Trp Pro Glu Glu Leu Glu Cys Asn Arg Leu Pro His 120 125 Cys Asp Asp Thr Val Pro Val Thr Ser His Pro His Thr Glu Leu Ser 135 140 Gly Pro Gln Lys Lys Ser Asp Gln Val Pro Arg Asp Ile Gly Phe Trp 150 155 Cys Pro Lys His Leu Arg Thr Ser Gly Asp Gln Gly Tyr Arg Phe Leu 170 165 Gly Ile Glu Gln Cys Ala Pro Pro Cys Pro Asn Met Tyr Phe Lys Ser 180 185

<210> 71 <211> 253 <212> PRT <213> Homo sapiens

Asp Glu Leu Asp Phe Ala 195

<400> 71

 Met
 Arg
 Asp
 Pro Gly
 Ala
 Ala
 Ala
 Pro Leu
 Ser
 Ser
 Leu Gly
 Leu Cys

 1
 5
 10
 15
 15

 Ala
 Leu Val
 Leu Ala
 Leu Leu Gly
 Ala
 Leu Ser
 Ala Gly
 Ala Gly
 Ala

 Ala
 Leu Val
 Bro
 Ala
 Bro
 Ala
 Bro
 Ala
 Ala
 Bro
 Ala
 Ala

90 Glu Leu Arg Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val 105 Leu Asp Gln Ala Ile Pro Pro Cys Arg Ser Leu Cys Glu Arg Ala Arg 120 Gln Gly Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln Trp Pro Glu 135 Arg Leu Arg Cys Glu Asn Phe Pro Val His Gly Ala Gly Glu Ile Cys 150 155 Val Gly Gln Asn Thr Ser Asp Gly Ser Gly Gly Pro Gly Gly Pro 170 175 165 Thr Ala Tyr Pro Thr Ala Pro Tyr Leu Pro Asp Leu Pro Phe Thr Ala 185 Leu Pro Pro Gly Ala Ser Asp Gly Arg Gly Arg Pro Ala Phe Pro Phe 205 200 Ser Cys Pro Arg Gln Leu Lys Val Pro Pro Tyr Leu Gly Tyr Arg Phe 215 Leu Gly Glu Arg Asp Cys Gly Ala Pro Cys Glu Pro Gly Arg Ala Asn 230 235 Gly Leu Met Tyr Phe Lys Glu Glu Glu Arg Arg Phe Ala 245

<210> 72 <211> 251

<212> PRT

<213> Mouse

<400> 72

Met Arg Gly Pro Gly Thr Ala Ala Ser His Ser Pro Leu Gly Leu Cys Ala Leu Val Leu Ala Leu Leu Gly Ala Leu Pro Thr Asp Thr Arg Ala 25 Gln Pro Tyr His Gly Glu Lys Gly Ile Ser Val Pro Asp His Gly Phe 40 Cys Gln Pro Ile Ser Ile Pro Leu Cys Thr Asp Ile Ala Tyr Asn Gln 55 Thr Ile Leu Pro Asn Leu Leu Gly His Thr Asn Gln Glu Asp Ala Gly 70 75 Leu Glu Val His Gln Phe Tyr Pro Leu Val Lys Val Gln Cys Ser Pro 85 90 Glu Leu Arg Phe Phe Leu Cys Ser Met Tyr Ala Pro Val Cys Thr Val 105 Leu Asp Gln Ala Ile Pro Pro Cys Arg Ser Leu Cys Glu Arg Ala Arg 120 Gln Gly Cys Glu Ala Leu Met Asn Lys Phe Gly Phe Gln Trp Pro Glu 135 140 Arg Leu Arg Cys Glu Asn Phe Pro Val His Gly Ala Gly Glu Ile Cys 155 Val Gly Gln Asn Thr Ser Asp Gly Ser Gly Gly Ala Gly Gly Ser Pro 165 170 Thr Ala Tyr Pro Thr Ala Pro Tyr Leu Pro Asp Pro Pro Phe Thr Ala 185 190 Met Ser Pro Ser Asp Gly Arg Gly Arg Leu Ser Phe Pro Phe Ser Cys 200 205 Pro Arg Gln Leu Lys Val Pro Pro Tyr Leu Gly Tyr Arg Phe Leu Gly 215 Glu Arg Asp Cys Gly Ala Pro Cys Glu Pro Gly Arg Ala Asn Gly Leu

Met Tyr Phe Lys Glu Glu Glu Arg Arg Phe Ala 245 <210> 73 <211> 277 <212> PRT <213> Homo sapiens <400> 73 Met Glu Trp Gly Tyr Leu Leu Glu Val Thr Ser Leu Leu Ala Ala Leu 10 Ala Leu Leu Gln Arg Ser Ser Gly Ala Ala Ala Ser Ala Lys Glu 25 Leu Ala Cys Gln Glu Ile Thr Val Pro Leu Cys Lys Gly Ile Gly Tyr 40 Asn Tyr Thr Tyr Met Pro Asn Gln Phe Asn His Asp Thr Gln Asp Glu 55 Ala Gly Leu Glu Val His Gln Phe Trp Pro Leu Val Glu Ile Gln Cys 70 Ser Pro Asp Leu Lys Phe Phe Leu Cys Ser Met Tyr Thr Pro Ile Cys 90 Leu Glu Asp Tyr Lys Lys Pro Leu Pro Pro Cys Arg Ser Val Cys Glu 105 Arg Ala Lys Ala Gly Cys Ala Pro Leu Met Arg Gln Tyr Gly Phe Ala 115 120 125 Trp Pro Asp Arg Met Arg Cys Asp Arg Leu Pro Glu Gln Gly Asn Pro 135 Asp Thr Leu Cys Met Asp Tyr Asn Arg Thr Asp Leu Thr Thr Ala Ala 150 155 Pro Ser Pro Pro Arg Arg Leu Pro Pro Pro Pro Gly Glu Gln Pro 165 170 Pro Ser Gly Ser Gly His Gly Arg Pro Pro Gly Ala Arg Pro Pro His 185 Arg Gly Gly Gly Gly Gly Gly Gly Asp Ala Ala Pro Pro 200 Ala Arg Gly Gly Gly Gly Gly Lys Ala Arg Pro Pro Gly Gly Gly 215 220 Ala Ala Pro Cys Glu Pro Gly Cys Gln Cys Arg Ala Pro Met Val Ser 235 230 Val Ser Ser Glu Arg His Pro Leu Tyr Asn Arg Val Lys Thr Gly Gln 245 250 Ile Ala Asn Cys Ala Leu Pro Cys His Asn Pro Phe Phe Ser Gln Asp 260 265 Glu Arg Ala Phe Thr 275 <210> 74 <211> 274 <212> PRT <213> Mouse <400> 74 Met Glu Trp Gly Tyr Leu Leu Glu Val Thr Ser Leu Leu Ala Ala Leu Ala Val Leu Gln Arg Ser Ser Gly Ala Ala Ala Ala Ser Ala Lys Glu 20

230

240



Leu Ala Cys Gln Glu Ile Thr Val Pro Leu Cys Lys Gly Ile Gly Tyr Asn Tyr Thr Tyr Met Pro Asn Gln Phe Asn His Asp Thr Gln Asp Glu Ala Gly Leu Glu Val His Gln Phe Trp Pro Leu Val Glu Ile Gln Cys 70 75 Ser Pro Asp Leu Lys Phe Phe Leu Cys Ser Met Tyr Thr Pro Ile Cys Leu Glu Asp Tyr Lys Lys Pro Leu Pro Pro Cys Arg Ser Val Cys Glu 105 Arg Ala Lys Ala Gly Cys Ala Pro Leu Met Arg Gln Tyr Gly Phe Ala 120 125 Trp Pro Asp Arg Met Arg Cys Asp Arg Leu Pro Glu Gln Gly Asn Pro 135 140 Asp Thr Leu Cys Met Asp Tyr Asn Arg Thr Asp Leu Thr Thr Ala Ala 150 155 Pro Ser Pro Pro Arg Arg Leu Pro Pro Pro Pro Pro Gly Glu Gln 170 165 Pro Pro Ser Gly Ser Gly His Ser Arg Pro Pro Gly Ala Arg Pro Pro 185 His Arg Gly Gly Ser Ser Arg Gly Ser Gly Asp Ala Ala Ala Pro 200 Pro Ser Arg Gly Gly Lys Ala Arg Pro Pro Gly Gly Ala Ala Pro 215 220 Cys Glu Pro Gly Cys Gln Cys Arg Ala Pro Met Val Ser Val Ser Ser 230 235 Glu Arg His Pro Leu Tyr Asn Arg Val Lys Thr Gly Gln Ile Ala Asn 250 Cys Ala Leu Pro Cys His Asn Pro Phe Phe Ser Gln Asp Glu Arg Ala 265 Phe Thr

<210> 75

<211> 231

<212> PRT

<213> Homo sapiens

<400> 75

Met Ala Val Ala Pro Leu Arg Gly Ala Leu Leu Trp Gln Leu Leu Ala Ala Gly Gly Ala Ala Leu Glu Ile Gly Arg Phe Asp Pro Glu Arg Gly Arg Gly Ala Ala Pro Cys Gln Ala Val Glu Ile Pro Met Cys Arg Gly Ile Gly Tyr Asn Leu Thr Arg Met Pro Asn Leu Leu Gly His Thr 55 Ser Gln Gly Glu Ala Ala Ala Glu Leu Ala Glu Phe Ala Pro Leu Val 75 Gln Tyr Gly Cys His Ser His Leu Arg Phe Phe Leu Cys Ser Leu Tyr 90 Ala Pro Met Cys Thr Asp Gln Val Ser Thr Pro Ile Pro Ala Cys Arg 105 100 Pro Met Cys Glu Gln Ala Arg Leu Arg Cys Ala Pro Ile Met Glu Gln 120 Phe Asn Phe Gly Trp Pro Asp Ser Leu Asp Cys Ala Arg Leu Pro Thr 140 130 135



Arg Asn Asp Pro His Ala Leu Cys Met Glu Ala Pro Glu Asn Ala Thr 155 Ala Gly Pro Ala Glu Pro His Lys Gly Leu Gly Met Leu Pro Val Ala 165 170 Pro Arg Pro Ala Arg Pro Pro Gly Asp Leu Gly Pro Gly Ala Gly Gly 185 Ser Gly Thr Cys Glu Asn Pro Glu Lys Phe Gln Tyr Val Glu Lys Ser 200 Arg Ser Cys Ala Pro Arg Cys Gly Pro Gly Val Glu Val Phe Trp Ser 215 Arg Arg Asp Lys Asp Phe Ala <210> 76 <211> 232 <212> PRT <213> Mouse <400> 76 Met Ala Val Pro Pro Leu Leu Arg Gly Ala Leu Leu Trp Gln Leu Leu Ala Thr Gly Gly Ala Ala Leu Glu Ile Gly Arg Phe Asp Pro Glu Arg Gly Arg Gly Pro Ala Pro Cys Gln Ala Met Glu Ile Pro Met Cys 40 Arg Gly Ile Gly Tyr Asn Leu Thr Arg Met Pro Asn Leu Leu Gly His Thr Ser Gln Gly Glu Ala Ala Ala Gln Leu Ala Glu Phe Ser Pro Leu Val Gln Tyr Gly Cys His Ser His Leu Arg Phe Phe Leu Cys Ser Leu 90 Tyr Ala Pro Met Cys Thr Asp Gln Val Ser Thr Pro Ile Pro Ala Cys 105 Arg Pro Met Cys Glu Gln Ala Arg Leu Arg Cys Ala Pro Ile Met Glu 120 125 Gln Phe Asn Phe Gly Trp Pro Asp Ser Leu Asp Cys Ala Arg Leu Pro 135 140 Thr Arg Asn Asp Pro His Ala Leu Cys Met Glu Ala Pro Glu Asn Ala 150 155 Thr Ala Gly Pro Thr Glu Pro His Lys Gly Leu Gly Met Leu Pro Val 170 Ala Pro Arg Pro Ala Arg Pro Pro Gly Asp Ser Ala Pro Gly Pro Gly 185 Ser Gly Gly Thr Cys Asp Asn Pro Glu Lys Phe Gln Tyr Val Glu Lys 200 205 Ser Arg Ser Cys Ala Pro Arg Cys Gly Pro Gly Val Glu Val Phe Trp 215 Ser Arg Arg Asp Lys Asp Phe Ala 230

<210> 77

<211> 227

<212> PRT

<213> Homo sapiens

<400> 77

Met Gln Arg Pro Gly Pro Arg Leu Trp Leu Val Leu Gln Val Met Gly

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Ser Cys Ala Ala Ile Ser Ser Met Asp Met Glu Arg Pro Gly Asp Gly
Lys Cys Gln Pro Ile Glu Ile Pro Met Cys Lys Asp Ile Gly Tyr Asn
                            40
Met Thr Arq Met Pro Asn Leu Met Gly His Glu Asn Gln Arg Glu Ala
                        55
Ala Ile Gln Leu His Glu Phe Ala Pro Leu Val Glu Tyr Gly Cys His
                    70
                                        75
Gly His Leu Arg Phe Phe Leu Cys Ser Leu Tyr Ala Pro Met Cys Thr
                                    90
               85
Glu Gln Val Ser Thr Pro Ile Pro Ala Cys Arg Val Met Cys Glu Gln
                                105
Ala Arg Leu Lys Cys Ser Pro Ile Met Glu Gln Phe Asn Phe Lys Trp
                            120
Pro Asp Ser Leu Asp Cys Arg Lys Leu Pro Asn Lys Asn Asp Pro Asn
                        135
Tyr Leu Cys Met Glu Ala Pro Asn Asn Gly Ser Asp Glu Pro Thr Arg
                    150
                                        155
Gly Ser Gly Leu Phe Pro Pro Leu Phe Arg Pro Gln Arg Pro His Ser
                165
                                    170
Ala Gln Glu His Pro Leu Lys Asp Gly Gly Pro Gly Arg Gly Gly Cys
                                185
Asp Asn Pro Gly Lys Phe His His Val Glu Lys Ser Ala Ser Cys Ala
                            200
                                                205
Pro Leu Cys Thr Pro Gly Val Asp Val Tyr Trp Ser Arg Glu Asp Lys
                        215
Arg Phe Ala
225
<210> 78
<211> 29
<212> PRT
<213> Homo sapiens
<400> 78
Asp Arg Val Val Cys Asn Asp Lys Phe Ala Glu Asp Gly Ala Arg Thr
Val Ala Gln Gly Thr Lys Lys Glu Gly Cys Thr Ile Leu
                                25
<210> 79
<211> 29
<212> PRT
<213> Mouse
<400> 79
Asp Arg Val Val Cys Asn Asp Lys Phe Ala Glu Asp Gly Ala Arg Thr
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Val Ala Gln Gly Thr Asn Lys Glu Gly Cys Thr Ile Leu
<210> 80
<211> 29
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<212> PRT

<213> Homo sapiens

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58

His Ala Ser Val Ala Cys Ser Arg Glu His Asn His Ile His Tyr Glu

<400> 85



15

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Thr Thr Gly Pro Ala Leu Cys Thr Ile Val
            20
<210> 86
<211> 30
<212> PRT
<213> Homo sapiens
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Asp Ser Thr Ala Cys Asn Lys Ala Asp Glu Lys Leu Glu Leu Gly Asp
                                    10
Thr Val Val Leu Gly Ser Gln Asn Lys Ala Cys Thr Val Leu
                                25
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<211> 30
<212> PRT
<213> Mouse
<400> 87
Asn Ser Thr Ala Cys Asn Lys Ala Asp Glu Lys Leu Glu Leu Gly Asp
                5
Thr Val Val Leu Gly Ser Lys Asn Lys Ala Cys Ser Val Val
                                25
<210> 88
<211> 29
<212> PRT
<213> Homo sapiens
<400> 88
Asp Arg Ala Val Cys Val Glu Arg Phe Ser Asp Asp Gly Tyr Arg Thr
                5
Val Ala Gln Gly Thr Lys Lys Glu Gly Cys Thr Ile Leu
                                25
<210> 89
<211> 29
<212> PRT
<213> Mouse
<400> 89
Asp Arg Ala Val Cys Val Glu Arg Phe Ser Asp Asp Gly Tyr Arg Thr
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Val Ala Gln Gly Thr Lys Lys Glu Gly Cys Thr Ile Leu
<210> 90
<211> 65
<212> PRT
<213> Homo sapiens
<400> 90
His Glu Lys Val Ala Cys Ser Gly Gly Ala Pro Gly Ala Gly Gly Ala
Gly Gly Ala Gly Gly Ala Ala Gly Ala Gly Ala Ala Gly Ala Gly
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Ala Gly Gly Pro Gly Gly Arg Gly Glu Tyr Glu Glu Leu Gly Ala Val Glu Gln His Val Arg Tyr Glu Thr Thr Gly Pro Ala Leu Cys Thr Val Val 65 <210> 91 <211> 66 <212> PRT <213> Mouse <400> 91 His Glu Lys Val Ala Cys Ser Gly Gly Ala Pro Gly Ala Gly Gly Arg Gly Gly Ala Gly Gly Ala Ala Ala Gly Ala Gly Ala Gly Arg Gly Ala Ser Ser Pro Gly Ala Arg Gly Glu Tyr Glu Glu Leu Gly Ala Val Glu Gln His Val Arg Tyr Glu Thr Thr Gly Pro Ala Leu Cys Thr Val Val 65 <210> 92 <211> 28 <212> PRT <213> Homo sapiens <400> 92 Ala Gln Ser Val Ala Cys Asp Gln Glu Ala Gly Ala Leu Tyr Val Ile Gln Glu Gly Leu Glu Asn Thr Gly Cys Thr Leu Val 20 <210> 93 <211> 28 <212> PRT <213> Mouse <400> 93 Ala Gln Ser Val Ala Cys Asp Gln Glu Ala Gly Ala Leu Tyr Val Ile Gln Glu Gly Leu Glu Asn Thr Gly Cys Thr Leu Val 20 <210> 94 <211> 28 <212> PRT <213> Homo sapiens <400> 94 Ala Glu Ser Ile Ala Cys Asp Arg Asp Ser Gly Gln Leu Tyr Val Ile Gln Glu Gly Leu Glu Ser Thr Gly Cys Thr Leu Val 20 25

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Gly Gln Val Asp Gly Asp Val Leu Ser Gly Val Cys Phe Val Gly Leu
Asn Asn Val Asp Ala Leu Arg Gly Phe
            20
<210> 96
<211> 25
<212> PRT
<213> Mouse
<400> 96
Gly Gln Val Asp Gly Asp Val Leu Ser Gly Val Cys Phe Leu Gly Leu
Asn Asn Val Asp Ala Leu Arg Gly Phe
            20
<210> 97
<211> 25
<212> PRT
<213> Homo sapiens
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Gly Gln Ile Asp Gly Asp Leu Leu Ser Gly Val Cys Phe Val Gly Leu
                                     10
Asn Ser Leu Asp Pro Leu Arg Gly Phe
            20
<210> 98
<211> 25
<212> PRT
<213> Homo sapiens
<400> 98
Asn Lys Ile Glu Gly Asp Asn Ile Ser Gly Val Cys Phe Val Gly Leu
                                     10
Tyr Asp Val Asp Ala Leu Arg Tyr Phe
            20
<210> 99
<211> 25
<212> PRT
<213> Mouse
<400> 99
Asn Lys Ile Glu Gly Asp Asn Ile Ser Gly Val Cys Phe Val Gly Leu
                                     10
                5
Tyr Asp Val Asp Ala Leu Arg Tyr Phe
            20
<210> 100
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<211> 25
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   <213> Homo sapiens
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   Arg Leu Val Asp Ala Asp Glu Leu Thr Gly Leu Cys Tyr Val Gly Asn
  Gln Asn Leu Asp Ala Leu Thr Gly Phe
               20
   <210> 101
  <211> 25
   <212> PRT
   <213> Mouse
  <400> 101
  Arg Leu Val Asp Ala Asp Glu Leu Thr Gly Leu Cys Tyr Val Gly Asn
                                       10
   Gln Asn Leu Asp Ala Leu Thr Gly Phe
               20
   <210> 102
   <211> 25
   <212> PRT
   <213> Homo sapiens
   <400> 102
  Ser Ser Val Asp Gly Asp Pro Val Ala Gly Ile Cys Tyr Val Gly Asn
  Gln Asn Leu Asn Ser Leu Arg Arg Phe
               20
   <210> 103
   <211> 25
   <212> PRT
   <213> Homo sapiens
  <400> 103
  Asn Lys Val Glu Gly Asp Asn Ile Ser Gly Val Cys Phe Val Gly Leu
                   5
  Tyr Asp Leu Asp Ala Ser Arg Tyr Phe
               20
   <210> 104
  <211> 25
  <212> PRT
  <213> Mouse
  <400> 104
  Asn Lys Val Glu Gly Asp Asn Ile Ser Gly Val Cys Phe Val Gly Leu
                   5
  Tyr Asp Leu Asp Ala Ser Arg Tyr Phe
               20
  <210> 105
  <211> 25
  <212> PRT
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<213> Homo sapiens
<400> 105
Gly Gln Val Asp Gly Asp Leu Leu Ser Gly Val Cys Tyr Val Gly Leu
                                    10
Ser Ser Val Asp Ala Leu Arg Gly Phe
<210> 106
<211> 25
<212> PRT
<213> Mouse
<400> 106
Gly Gln Val Asp Gly Asp Leu Leu Ser Gly Val Cys Tyr Val Gly Leu
                5
                                    10
Ser Ser Val Asp Ala Leu Arg Gly Phe
            20
<210> 107
<211> 25
<212> PRT
<213> Homo sapiens
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Ser Ser Val Asp Gly Asp Pro Val Ala Gly Ile Cys Tyr Val Gly Asn
                                    10
Gln Ser Leu Asp Asn Leu Arg Gly Phe
            20
<210> 108
<211> 25
<212> PRT
<213> Mouse
<400> 108
Ser Ser Val Asp Gly Asp Pro Val Ala Gly Ile Cys Tyr Val Gly Asn
                5
                                    10
Gln Ser Leu Asp Asn Leu Arg Gly Phe
           20
<210> 109
<211> 25
<212> PRT
<213> Homo sapiens
<400> 109
Arg Lys Val Ala Gly Asp Glu Leu Thr Gly Leu Cys Tyr Val Ala Ser
                5
                                    10
Thr Asp Ala Ala Leu Thr Gly Phe
            20
<210> 110
<211> 25
<212> PRT
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A

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<210> 115 <211> 32 <212> PRT

<213> Homo sapiens <400> 115

Ala Tyr Arg Gly Ile Trp Glu Thr Thr Trp Ile Gln Glu Arg Cys Arg Glu Tyr His Ile Pro Cys Pro Tyr Gln Val Thr Gln Met Ser Arg Pro

<210> 116 <211> 32

<212> PRT <213> Mouse

<400> 116

Ala Tyr Arg Gly Ile Trp Glu Thr Trp Ile Gln Glu Arg Cys Arg 10 Glu Tyr His Ile Pro Cys Pro Tyr Gln Val Thr Gln Met Ser Arg Pro 20 25

<210> 117 <211> 17 <212> PRT

<213> Homo sapiens

<400> 117

Ser Asn Trp Ala Leu Phe Arg Tyr Ser Ala Asp Asp Ser Asn Met Ala 5 ' Val

<210> 118 <211> 17 <212> PRT <213> Mouse

<400> 118

Ser Asn Trp Ala Leu Phe Arg Tyr Ser Ala Asp Asp Ser Asn Met Ala Val

<210> 119 <211> 26 <212> PRT <213> Homo sapiens

His Tyr Arg Glu Ser Trp Glu Ala Ala Leu Thr Cys Ala Cys Pro Gly 5

His Asp Thr Gly Gln Pro Arg Ala Lys Pro 20

<210> 120 <211> 32 <212> PRT

<400> 119

<213> Homo sapiens

<400> 120 Val Asn Arg Ile Thr Trp Glu Ile Thr Trp Val Ser Asp His Cys Arg Gln Tyr His Ile Pro Cys Pro Tyr Gln Ala Lys Ala Lys Ala Arg Pro 20 25 <210> 121 <211> 32 <212> PRT <213> Mouse <400> 121 Val Asn Arg Ile Thr Trp Glu Met Thr Trp Phe Ser Asp His Cys His 10 Gln Tyr Arg Ile Pro Cys Pro Tyr Gln Ala Asn Pro Lys Ala Arg Pro 20 25 <210> 122 <211> 32 <212> PRT <213> Homo sapiens <400> 122 Ala Phe Arg Glu His Trp Glu Arg Thr Trp Leu Leu Gln Thr Cys Lys 10 Ser Tyr Ala Val Pro Cys Pro Pro Gly His Phe Pro Pro Met Ser Pro 25 <210> 123 <211> 32 <212> PRT <213> Mouse <400> 123 Ala Phe Arg Glu His Trp Glu Arg Thr Trp Leu Leu Gln Thr Cys Lys 10 Ser Tyr Ala Val Pro Cys Pro Pro Arg His Phe Ser Pro Met Ser Pro 25 30 <210> 124 <211> 26 <212> PRT <213> Homo sapiens <400> 124 His Asn Arg Pro Arg Trp Glu Ala Thr His Asn Cys Pro Cys Leu Arg 5

Asp Leu Gln Pro Asp Gln Ala Arg Arg Pro

<210> 125 <211> 26 <212> PRT

<213> Mouse

<400> 125

His Asn Arg Pro Arg Trp Glu Ala Thr His Asn Cys Pro Cys Leu Arg

15 Asp Leu Gln Pro Asp Gln Ala Arg Arg Pro 20 <210> 126 <211> 35 <212> PRT <213> Homo sapiens <400> 126 Leu Asn Met Asp Phe Trp Arg Leu Arg Ala Thr Glu Gln Pro Cys Ala 5 10 Ala Ala Gly Pro Gly Gly Arg Arg Asp Cys Ser Leu Pro Gly Gly 25 Ser Val Pro 35 <210> 127 <211> 35 <212> PRT <213> Mouse <400> 127 Leu Asn Met Asp Phe Trp Arg Leu Arg Ala Thr Glu Gln Pro Cys Thr 10 Ala Ala Thr Val Pro Gly Gly Arg Arg Asp Cys Ser Leu Pro Gly Gly 25 Ser Val Pro 35 <210> 128 <211> 33 <212> PRT <213> Homo sapiens <400> 128 Leu Asn Met Asp Tyr Trp Lys Ile Leu Ala Ala Gln His Lys Cys Lys Met Asn Asn Gln Thr Lys Thr Leu Asp Cys Leu Met Ala Ala Ser Ile 20 25 Pro <210> 129 <211> 48 <212> PRT <213> Human <400> 129 Val Gly Gln Asn Thr Ser Asp Lys Gly Thr Pro Ser Leu Leu Pro Glu Phe Trp Thr Ser Asn Pro Gln His Gly Gly His Arg Gly Gly Phe 25 Pro Gly Gly Ala Gly Ala Ser Glu Arg Gly Lys Phe Ser Cys Pro Arg 40

67

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Ala Pro Pro Pro Gly Leu Gln Pro Gly Ala Gly Gly Thr Pro Gly Gly
Pro Gly Gly Gly Ala Pro Pro Arg Tyr Ala Thr Leu Glu His Pro
                          40
        35
Phe His Cys
   50
<210> 131
<211> 26
<212> PRT
<213> Human
<400> 131
Leu Val Asp Leu Asn Leu Ala Gly Glu Pro Thr Glu Gly Ala Pro Val
                5
Ala Val Gln Arg Asp Tyr Gly Phe Trp Cys
            20
<210> 132
<211> 20
<212> PRT
<213> Human
<400> 132
Cys Met Glu Gly Pro Gly Asp Glu Glu Val Pro Leu Pro His Lys Thr
Pro Ile Gln Pro
            20
<210> 133
<211> 46
<212> PRT
<213> Human
<400> 133
Cys Met Asp Tyr Asn Arg Ser Glu Ala Thr Thr Ala Pro Pro Arg Pro
Phe Pro Ala Lys Pro Thr Leu Pro Gly Pro Pro Gly Ala Pro Ala Ser
                                25
Gly Gly Glu Cys Pro Ala Gly Gly Pro Phe Val Cys Lys Cys
<210> 134
<211> 26
<212> PRT
<213> Human
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10

Thr Phe Asp Pro His Thr Glu Phe Leu Gly Pro Gln Lys Lys Thr Glu

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